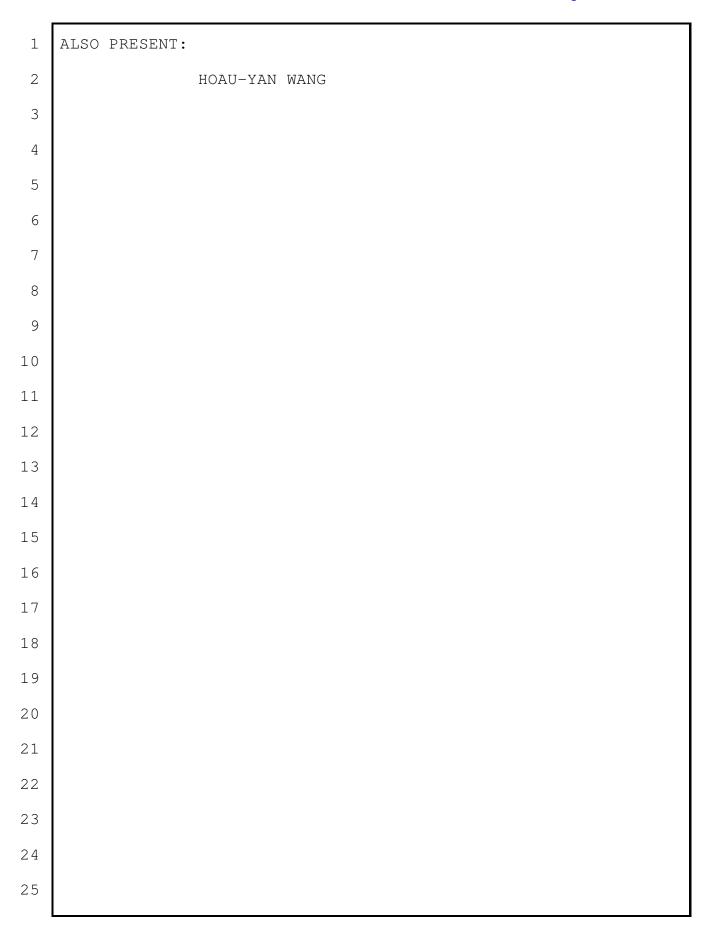
EXHIBIT 3

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IN THE UNITED STATES DISTRICT COURT
 1
                      FOR THE DISTRICT OF MARYLAND
                            SOUTHERN DIVISION
 2
 3
    UNITED STATES OF AMERICA
 4
         Plaintiff,
 5
                                  ) Case Number
              VS.
                                  )8:24-cr-00211-TDC-1
    HOAU-YAN WANG
 6
 7
         Defendant.
 8
                          TRANSCRIPT OF MOTIONS HEARING
 9
                     BEFORE THE HONORABLE THEODORE D. CHUANG
                           UNITED STATE DISTRICT JUDGE
                    TUESDAY, SEPTEMBER 30, 2025 at 9:04 a.m.
10
11
    APPEARANCES:
12
    On Behalf of the Plaintiff:
13
             UNITED STATES ATTORNEY'S OFFICE - DOJ
                   ANDREW TYLER, ESQUIRE
14
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                   VASANTH RAMAN SRIDHARAN, ESQUIRE
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16
    On Behalf of the Defendant:
17
18
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22
                        KIMBERLY A. BURSNER, RPR
                    FEDERAL OFFICIAL COURT REPORTER
                     6500 CHERRYWOOD LANE, SUITE 200
23
                        GREENBELT, MARYLAND 20770
24
                              (301) 344-3499
25
         ***COMPUTER AIDED TRANSCRIPTION OF STENOTYPE NOTES***
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DEPUTY CLERK:
                             All rise.
                                        The United States District
 1
    Court for the District of Maryland is now in session.
 2
    Honorable Theodore D. Chuang presiding.
 3
                          Thank you. Please be seated.
              THE COURT:
 4
 5
              DEPUTY CLERK:
                             The matter now pending before this
    Court is Criminal Action Number 24-0211-TDC, United States of
 6
 7
    America versus Hoau-Yan Wang.
 8
         We are here today for the purpose of a Daubert hearing.
 9
         Counsel, please identify yourselves, for the record.
10
                         Good morning, Your Honor. Andrew Tyler
11
    for the United States. I'm joined here at counsel table also
12
    by our paralegal, Anna Cornich, and I'll let my colleagues
13
    introduce themselves.
14
              MR. SRIDHARAN: Good morning, Your Honor. Vasanth
15
    Sridharan on behalf of the United States.
16
              MR. PATHAN: Good morning, Your Honor. Kashan Pathan
17
    on behalf of the United States.
18
              THE COURT:
                          Okay. Good morning, everyone.
19
              MS. BEIDEL: Good morning, Your Honor. Jennifer
20
    Beidel and Emma Blackwood on behalf of Dr. Hoau-Yan Wang.
21
              THE COURT:
                          Good morning. Good morning, Dr. Wang.
22
         So we are here for a hearing on basically on the motions
23
    regarding -- well, the motions in limine regarding expert
24
    witnesses, primarily the one regarding Dr. Brookes which I
25
    think is sort of the gateway to the other motions.
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I've reviewed the briefs and the exhibits.
                                                 I do have some
questions. I, frankly, think this is the kind of issue that
requires me to understand exactly what this witness would
testify to both as qualifications and his opinions and what the
counter-arguments are.
     As you know and the case law sometimes in a civil case we
can do that through deposition transcript review, but this is
not a civil case, so are we prepared to have Dr. Brookes
testify today?
                     Yes, Your Honor. We prepared pretty
          MR. TYLER:
extensive testimony for Dr. Brookes today. Also just to
apprise the Court, he does have a very hard stop at 2:30 this
afternoon because he's chairing a thesis defense that he's
arranged to do remotely, but still --
                     Can you just speak into the microphone?
          THE COURT:
                     I'm just reminding the Court that Dr.
          MR. TYLER:
Brookes has a 2:30 hard stop this afternoon because he's
chairing a thesis defense which he's arranged to do remotely,
but, obviously, will need to be off the stand to do that at
that time.
          THE COURT: I didn't know that before, but, honestly,
I didn't -- I was hoping it would not take that long, so I will
see how that goes. I think I have a sentencing hearing at 2:30
myself. Let's just get going and see how it goes.
```

Is there anything we should discuss, though, before the

testimony from either side? 1 Not from the government, Your Honor. 2 MR. TYLER: MS. BEIDEL: Not from the defense either, Your Honor. 3 THE COURT: So why don't we have him come forward. 4 5 GOVERNMENT TESTIMONY 6 7 PAUL BROOKES, after having been duly 8 sworn, was examined and testified as follows: 9 And please spell your first and last 10 DEPUTY CLERK: 11 name, for the record. 12 THE WITNESS: P-a-u-l B-r-o-o-k-e-s. DEPUTY CLERK: 13 Thank you, sir. THE COURT: Go ahead. 14 15 DIRECT EXAMINATION BY MR. TYLER: 16 17 Good morning, Dr. Brookes. Good morning. 18 19 Could you tell us your title of witness? 20 I'm a tenured professor of anesthesiology at the University of Rochester Medical Center. 21 22 And could you give us a little bit about your educational 23 background? Yes. I received my PhD at University College, London, in 24 25 biochemistry in 1993. I then went to Cambridge University in

- 1 the U.K. and got a PhD also in biochemistry in 1997. I, then,
- 2 did a post-doctoral fellowship at the Institute of Neurology in
- 3 University of London for a year and a half before moving to the
- 4 U.S.
- 5 Q. With respect to your time at undergrad, did you do any
- 6 | work with respect to neuroscience during that time?
- 7 A. Yes. I was a lab technician in the lab in the Institute
- 8 of Neurology which is connected with University College London.
- 9 That's the lab that I returned to as a post-op fellow.
- 10 **Q.** And did your doctoral training include statistical
- 11 | training?
- 12 A. Yes, it did.
- MR. TYLER: Ms. Cornich, if you could pull up
- 14 Exhibit-34, please.
- 15 BY MR. TYLER:
- 16 Q. Dr. Brookes, could you tell us what this is?
- 17 **A.** This is my academic CV.
- 18 Q. And does that CV contain a lot of information about your
- 19 | society memberships, awards, N.I.H. grants, reviewed
- 20 publications, conference, abstracts, et cetera?
- 21 A. Correct.
- 22 MR. TYLER: At this time the government would like to
- 23 move into evidence Exhibit-34.
- 24 | THE COURT: Okay. Exhibit-34 is in evidence. I
- 25 | think, am I correct, the binders I have correspond to what you

```
are referring to?
 1
                          Yes, Your Honor.
 2
              MR. TYLER:
              THE COURT:
                          Do you know whether any of the exhibits
 3
    will have objections?
 4
 5
              MR. TYLER: I do not, Your Honor.
                          Well, just to kind of be efficient, as
              THE COURT:
 6
 7
    you know from my procedures, I want to make sure all the
 8
    exhibits are offered and admitted so the record is clear, but
 9
    ideally, if there is no objections, we can kind of move more
10
    swiftly and maybe even do them in groups, so I'm counting on,
11
    Ms. Beidel, if you could just jump up right away if there is an
12
    issue, or when there is an opportunity to confer of what is
13
    objected to and what is not.
14
         So Exhibit-34 is in evidence.
15
              MR. TYLER:
                          Thank you, Your Honor.
         Ms. Cornich, could you also bring up Exhibit-35?
16
17
    BY MR. TYLER:
         And, Dr. Brookes, can you tell us what this is?
18
19
         This is a shorter version of my CV focusing my work in the
20
    area of research integrity.
21
              MR. TYLER:
                         At this time, the government also move
22
    into evidence Exhibit-35.
23
              THE COURT: Okay. Exhibit-35 is in evidence for
24
    purposes of the hearing.
    BY MR. TYLER:
25
```

- 1 Q. Dr. Brookes, you mentioned your time at the Institute of
- 2 Neurology. What did you work on while you were there?
- 3 **A.** I was working in a lab studying mitochondria in the
- 4 | context of Parkinson's and Alzheimer's Disease.
- 5 Q. When you came to the United States, where did you come and
- 6 what did you do there?
- 7 A. I moved to the University of Alabama at Birmingham, also
- 8 working on mitochondria, but this time in the context of
- 9 cardiovascular disease.
- 10 **Q.** During that time, did you do work with Western blots?
- 11 A. Correct. UAB was where I was trained in the technique of
- 12 Western blots and regularly used that techniques.
- 13 **Q.** Approximately, how many did you do per year during your
- 14 | time there?
- 15 **A.** I would estimate roughly a hundred Western blots a year,
- 16 so a couple a week.
- 17 \mathbf{Q} . Are you familiar with a technique called blue native gels?
- 18 | A. Correct. Blue native gel is a specific type of gel for
- 19 isolating protein complexes in their native format. And, in
- 20 | fact, we developed a technique while I was at UAB in order to
- 21 | be able to Western blot that type of gel which nobody had done
- 22 before.
- 23 **Q.** Are you published on that topic?
- 24 | A. Correct. That is in a paper in the Journal of Proteomics
- 25 | in 1992.

- 1 Q. After you left the University of Alabama Birmingham, where
- 2 did you go from there?
- 3 **A.** From UAB, I was recruited to the University of Rochester
- 4 in August of 20803.
- 5 **Q.** And did you bring any funding with you when you came to
- 6 | the University of Rochester?
- 7 A. Yes. While I was a post-doc at UAB, I wrote and submitted
- 8 N.I.H. RO1 grant that was successfully funded, so then when I
- 9 arrived in Rochester, that was the funding that I brought with
- 10 me to start my lab.
- 11 **Q.** And, generally speaking, what experience have you had with
- 12 N.I.H. grants since then?
- 13 A. So, that grant that was awarded in 2003. I still have
- 14 that grant to this day. It has been continuously funded for
- 15 over 20 years. It's currently on its fifth funding cycle. In
- 16 addition, I've had another four RO1 grants as a principal
- 17 | investigator, and I've been a coinvestigator on a number of
- 18 other grants from N.I.H., as well as other funding bodies.
- 19 Q. During your time at the University of Rochester, have you
- 20 | conducted Western blot experiments there?
- 21 | A. Yes. My lab still uses Western blotting to this day. I
- 22 | would say anywhere between one and five per week depending on
- 23 | what project we're working on. Again, averaging out to about a
- 24 hundred a year.
- 25 \mathbf{Q} . And do you do that using film still development or digital

development?

- 2 A. Up until right before COVID, we used film development for
- 3 Western blotting and then right before COVID as the technology
- 4 became cheaper, we purchased a digital imaging system to
- 5 quantify those blots.
- 6 Q. Dr. Brookes, do you also teach at the university?
- 7 **A.** Correct. I teach in the medical school and the graduate
- 8 school.
- 9 Q. And do you teach any courses relating to ethics?
- 10 **A.** Yes. So there is a mandatory ethics course for all
- 11 | incoming medical and graduate students and I teach one of the
- 12 | modules in that course.
- 13 **Q.** And during the course of your career, have you had any
- 14 | experience consulting with private companies who are attempting
- 15 to develop drugs?
- 16 A. Yes. I have consulted in on the advisory boards of a
- 17 | number of early stage pharmaceutical companies. I've had
- 18 | companies send drug material to my lab for experimentation via
- 19 | material transfer agreements, as well as other types of
- 20 | consulting work.
- 21 **Q.** Turning to image analysis specifically, how did you first
- 22 start doing that?
- 23 | A. I was reviewing a grant for the American Heart Association
- 24 | and I found, as is frequently the case when you are reviewing
- 25 grants in PDF format, you may use a split screen to view

1 different parts of the grant at the same time. And I noticed

- 2 that two of the images in the grant were duplicated to
- 3 represent different proteins. The Western blot specifically
- 4 had been reused and flipped around. I reported that to the
- 5 AHA. I also forwarded my findings to the U.S. Office of
- 6 Research Integrity. They subsequently performed an
- 7 | investigation and made a finding of misconduct against the
- 8 author of that grant.
- 9 **Q.** Approximately, when was that?
- 10 **A.** That was late 2010, early 2011.
- 11 **Q.** And then what happened after that with respect to your
- 12 | work on image analysis?
- 13 | A. I continued investigating any and all grants and papers
- 14 | that came across my desk. I did, in fact, find some suspicious
- 15 | images in papers from colleagues at my own university. I
- 16 reported those to my university and not much happened in terms
- 17 of action.
- 18 At the time there were a series of public facing blogs
- 19 where people would report on these types of issues with the
- 20 | images and live sciences papers. And so in 2012, I decided to
- 21 join in and start an anonymous blog.
- 22 **Q.** What work did you do for that blog?
- 23 | A. I found and reported on problem images in papers. Over
- 24 | the course of the lifetime of that blog, we reported on about
- 25 | 300 papers. That resulted in several retractions and a number

1 of other outcomes for the authors of those papers.

- Q. At some point, did there come a time when you -- that blog was de-anonymized?
- 4 A. Yes. That was an anonymous blog. At the time I was a
- 5 fellow junior faculty member without tenure, so I considered it
- 6 | necessary to remain anonymous. However, once my identity was
- 7 | compromised, my university essentially got involved. They
- 8 essentially told me to shut down the blog and as somebody
- 9 | without tenure, I didn't have much choice in the matter. I was
- 10 | also told that my activities in that area were separate from my
- 11 university employment and, therefore, the university would not
- 12 be willing to provide me with any legal support. So I was
- 13 advised to obtain a lawyer and that lawyer was able to rebut
- 14 numerous attempts to sue me that were made.
- 15 **Q.** And then after that time working in your personal
- 16 | capacity, did you continue to do work in image analysis?
- 17 | A. Yes. I still do to this day. I have a master database of
- 18 | several thousand examples of image analysis and image problems.
- 19 | I continue to post using my real identity, my real name on a
- 20 | public website known as Pub Peer. That is a website sort of a
- 21 clearinghouse for issues in the bioscience literature. My work
- 22 has led to hundreds of retractions of papers. In addition, my
- 23 | collated databases and annotated databases and taxonomy of
- 24 | image problems has been used by collaborators to train
- 25 | machine-loading algorithms and AI. As you may be aware, there

- 1 are a number of AI detection tools for this type of image
- 2 problems nowadays.
- 3 Q. Do journals consult with you for purposes of viewing that
- 4 | image analysis?
- 5 **A.** Yes. I am regularly requested to look over papers for
- 6 which editors may be suspicious of problems in the data.
- 7 Q. Over the years, have you become familiar with various
- 8 tools that you used in this case to do analysis?
- 9 A. Correct. Yes.
- 10 **Q.** And that includes various PowerPoint Photoshop and image J
- 11 tools?
- 12 **A.** Yes.
- 13 **Q.** Are you familiar with published literature about image
- 14 | analysis and research misconduct?
- 15 A. Correct. Yes. And I have published in that area as well.
- 16 **Q.** And are you familiar with something known as the JCB
- 17 | standards?
- 18 A. Correct. So that refers to a set of standards that were
- 19 | first captured in an editorial in the Journal of Cell Biology
- 20 by the editor and chief at the time Mike Rossner and that
- 21 essentially describes what you are and are not allowed to do
- 22 | with a scientific image when preparing it for publication. Do
- 23 | you want me to go in to the details of that now or --
- 24 | Q. Yes. So what are those standards in the --
- 25 **A.** So there are a couple of core concepts to that editorial.

1 One is that any manipulation of the scientific image should not

- 2 change the informational content of the image and then the
- 3 other principle is if an adjustment is made to an image, such
- 4 as brightness or contrast, the whole image must be adjusted
- 5 evenly across the field of view. It is not acceptable to
- 6 brighten or darken particular features of an image.
- 7 **Q.** Why is it not okay to brighten or darken one image or one
- 8 lane? Sorry. One lane?
- 9 A. Because that would change the informational content of the
- 10 image.
- 11 **Q.** And is that informational content equivalent to the data
- 12 of the experiment?
- 13 **A.** Sorry?
- 14 **Q.** Is the informational content equivalent to the data of the
- 15 | experiment?
- 16 A. Yes. In the case of Western blotting, for example, the
- 17 darkness of the bands in the Western blot is the data. That is
- 18 | the information contained within the image, but is then
- 19 subsequently used to draw a conclusion.
- 20 **Q.** You mentioned AI learning a minute ago. Do there have to
- 21 be sufficient standards in order to make that type of analysis
- 22 | codifiable?
- MS. BEIDEL: Objection, Your Honor. I haven't been
- 24 | objecting to the leading for purpose of facilitating this
- 25 | hearing, but there is quite a lot of it and we are certainly

more interested in hearing Dr. Brookes' recitation of these
facts than Mr. Tyler's.

THE COURT: Okay. I understand the point and we should be careful about that. Why don't you reask the question?

BY MR. TYLER:

- Q. Dr. Brookes, with respect to AI software, how does that work in terms of in order to be able to code it for software?
- A. So, first of all, you know, for AI in order to be able to detect image problems, it needs a training set, but that training set is useless unless taxonomy has been applied to classify those examples of image manipulations. That taxonomy could include things like has the image been flipped, has it been darkened, has it been lightened, has it been recolored, has it been resized, have more than one image been overlaid on top of each other.

And so all of that information effectively is what is known as codifiable. It means that the standards and what has been done to the image can be distilled down into a set of instructions and, therefore, a computer can learn that.

- Q. And you mentioned that you have been published on this topic. Can you explain what you mean by that?
- A. So one of the papers that I published in that area, that
 is actually a preprint on bio-archive, which is a preprint
 server, that has not yet published, but that is a collaboration

with a group that is developing machine-loading algorithms.
Daniel Acuna and Konrad Kording.

The other paper is based on my experiences with the blog which was essentially investigating the impact of public discussion of problems in the bioscience literature on the degree of action that is taken subsequently about those papers.

And the conclusion of that paper was public discussion of problematic images in scientific papers results in between four and seven fault greater levels of action. By action, I mean, retractions, corrections and other things that journals do in response to such allegations.

The third paper was published last year. That is essentially an overview of the various tools available for doing this type of image analysis. That was essentially a conference proceeding. I was invited to a research integrity conference at the University of Pennsylvania, and all the speakers were invited to contribute a paper. As it ended up, all being published in a special issue of a journal.

- Q. Have you also been invited to speak on this topic?
- A. Yes. The conference proceeding that I just mentioned is one. I also participated in another conference during COVID on-line in another panel and then, in fact, this fall I'm due to sit on another panel at the annual meeting of the Association of Research Integrity Offices, ARIO.
 - Q. You mentioned hundreds of retractions. Have there been

- any instances where you discovered you made a mistake? 1
- Everybody makes mistakes. One particular example is 2
- I did call out an image on-line from a colleague at the 3
- 4 University of Rochester. He came angrily storming in to my
- 5 office and slammed down the original data on my desk and was
- able to prove that it was not, in fact, manipulated. 6
- 7 profusely apologized and we both moved on with our careers.
- 8 Do you consider that the information that you evaluated in
- 9 this case to be analogous to that?
- 10 Could you repeat the question, please?
- 11 MS. BEIDEL: Objection.
- 12 THE COURT: Why don't you rephrase.
- 13 BY MR. TYLER:
- 14 Dr. Brookes, do you consider the amount of information you
- 15 have for it to do the work in this case analogous to the amount
- of information you had in that instance? 16
- 17 Specifically, generally, when investigating these types of
- image issues in the literature, all you have is the published 18
- 19 figure. So it can actually be difficult to draw conclusions.
- However, in this case, I was afforded access to significantly 20
- 21 larger amounts of data including some materials that we use to
- 22 prepare the final published figures. So this enabled us to
- draw more certain conclusion. 23
- 24 MR. TYLER: Your Honor, may I have a movement to
- 25 consult with defense counsel about exhibits?

1 THE COURT: Sure.

2 MR. TYLER: Your Honor, may I approach the witness?

THE COURT: Yes. Does he have these binders or not

or anyone like that or --

have him verify.

3

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5 MR. TYLER: Your Honor, I have electronic versions 6 that he initialed yesterday which I was going to approach and

8 **THE COURT:** Okay. That's fine.

MS. BEIDEL: Your Honor, just to confirm --

THE COURT: Just pull the microphone up.

11 MS. BEIDEL: My apologies. We don't have objections

12 to any of the exhibits, except two that we are reserving on. I

13 think Mr. Tyler will tell you about that in a second, but just

14 | for purposes of trial, we may have different objections that I

15 | want to preserve.

16 THE COURT: No. I understand. I think we should

17 | just use this for use of the hearing. There could be different

18 issues at the trial. I understand that.

19 MS. BEIDEL: Thank you, Your Honor.

20 BY MR. TYLER:

- 21 **Q.** Dr. Brookes, do you recognize those disks in front of you?
- 22 **A.** Yes. Those are electronic materials that I reviewed
- 23 | yesterday and then subsequently initialed and dated.
- 24 **Q.** Generally speaking, what are the contents of those
- 25 exhibits?

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- They consist of PDF copies, as well as some Excel and 1 Α.
- PowerPoint files. The PDFs are the grants and papers, as well 2
- as reports that I submitted. 3
- At this time, the government moves in to 4 MR. TYLER:
- 5 evidence Exhibits-1 through 49.
- THE COURT: Exhibits-1 through 49 are in evidence for 6
- 7 the hearing.
- BY MR. TYLER: 8
- 9 Now, Dr. Brookes, turning to Western blots. Could you
- 10 tell us what the purpose of a Western blot is?
- 11 A Western blot serves to measure the relative amounts of
- 12 protein in one or more biological samples.
- 13 And how is it measured? What is it measured in?
- 14 So in a Western blot, you're separating proteins and then
- 15 exposing them to antibodies. The antibodies specifically
- recognize the protein. The antibodies then subject to a 16
- 17 chemical reaction that gives off light, so what you are
- measuring in a Western blot is the amount of light given off 18
- which is proportional to the amount of protein in a sample. 19
- 20 And that light is captured by a piece of x-ray film so the
- 21 darkness of the film correlates to the amount of protein in the
- 22 gel.
- 23 MR. TYLER: Your Honor, may I approach the witness
- 24 again?
- 25 THE COURT: Okay.

BY MR. TYLER:

- 2 Q. Dr. Brookes, do you see an envelope with an exhibit
- 3 labeled 52?

- 4 A. Correct. Fifty-two. Yes.
- 5 Q. Can you tell us what is in there?
- 6 A. These are some of the equipment and materials that are
- 7 typically used in the Western blotting process.
- 8 Q. Could you describe that using those tools and actually
- 9 demonstrate how a Western blot is done?
- 10 **A.** Yes. Okay. So, the first step is we have a pair of glass
- 11 plates that are separated by a spacer of 1.5 millimeters. We
- 12 | fill the gap in between these two plates with a solution that
- 13 then sets that is a gel. So it's just like Jello that you
- 14 | would eat at home. Right before it sets, we insert a small
- 15 thing called a comb in to the gel. Once the gel is set, we
- 16 remove the comb and that leaves behind a series of indentations
- 17 | in the well which are known as wells.
- 18 The protein is then -- the protein samples are then loaded
- 19 | in to those wells. We typically leave one well empty in order
- 20 to add a series of molecular weight markers in one of the
- 21 lanes. An electric current is then applied across the gel and
- 22 the proteins in this case are all coated with a molecule called
- 23 | SDS. That's sodium dodecyl sulfate. That's a detergent. That
- 24 gives all the proteins a negative charge so they might
- 25 | breakdown the gel towards the positive electrode. The proteins

are separated on the basis of their size.

So the small proteins migrate fast and end up at the bottom of the gel. Whereas, the larger proteins are retained at the top of the gel because they migrate more slowly. So you can think of the gel as sort of a size siv. When the gel is finished, we take the plate apart. We then put the gel in another apparatus with a membrane. The membrane is made up of paper called nitrocellulose which binds protein. We then transfer the gel to this membrane. So that transfer process is actually the blot. That is the process of Western blotting is transferring the gel to the membrane. The reason we do that is the gel is kind of floppy and fragile. The membrane is a piece of paper which is robust which we can then do things do.

So the membrane is then subjected to a series of washing steps and solutions containing antibodies and detection molecules. You can see on the side of this gel, perhaps you can see red and blue. Those are the molecular weight markers in one of the lanes. And then we add a chemical which is going to react with the antibody to give off light. And then we place the membrane inside a cassette. This is a film cassette, an x-ray film cassette.

So the membrane is placed in the film cassette in to the cassette in a dark room is introduced a piece of x-ray film.

This is a blank piece of old film. The x-ray films goes in to the cassette. The lid is closed for a set amount of time. Ter

seconds, 30 seconds, two minutes, five minutes.

After the given exposure, we take out the x-ray film and we put it through a developer machine, and it develops the Western blot, and so we get this type of image.

So this is a developed Western blot. The dark --

THE COURT: Can you point it towards me, would you mind?

THE WITNESS: Sorry. The dark things are bands. The dark things are bands and then alongside the bands we also annotate the different molecular weight markers. The way we do that is while we develop this x-ray film, the membrane stays in the cassette so then when we have the developed film, we can put this back into the cassette and transpose the molecular weight markers and other details about the experiment onto the film.

And so this developed film where you can see each -- there is a date. The corner of the film is folded so we know which was the top corner so we can orient it correctly. The details of the antibody are written in the side and molecular weight markers are listed down both sides of the gel.

So that's what we keep in a folder in the lab. That is the Western blot in the developed form. That is the data.

- Q. And then once you have that, what is the next step?
- A. So, typically, we have a flat bad scanner in the lab. And so we would take a digital scan of this piece of film, and then

we use a software called Image J. That is a free software 1 distributed by the N.I.H. and that is used to perform a process 2 3 called densitometry. So densitometry is essentially determining the darkness of the bands in each lane which allows 4 5 us to then get quantitative data out of this type of gel to make a graph. 6 7 And, Dr. Brookes, are you also familiar with something 8 called an isoelectric focusing Western blot? 9 Yes. So, the technique that I just described that was 10 called SDS Page. SDS, the detergent, and then polyacrylamide 11 gel electrophoresis. That gives everything a negative charge 12 so the proteins all move in one direction, so they are 13 separated by their molecular weight. If we do the gel slightly 14 differently, there is a technique called isoelectric focusing. 15 So we leave the SDS out and this time even without the detergent, all proteins have a negative charge. Proteins are 16 17 covered with positive and negative charges. Every protein is different. 18 19 And so we put the proteins in the gel on what is called a 20 pH gradient, so the pH ten is at the top. PH three is at the bottom. And when we apply an electric current, the proteins 21 22 will move up and down the gel until they reach their happy 23 place, a point where they don't have a net charge. And that 24 point is called the isoelectric point. So we shorthand that to

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PI.

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So the isoelectric point, in effect, is the pH value at which a protein no longer has a net charge. And, again, that is different for all types of proteins.

The gel process is different. Once an isoelectric focusing gel is complete, everything we described in terms of taking that gel, imprinting it onto a membrane, putting the membrane in the cassette with the film, putting the markers on the piece of developed film, that whole downstream process is the same.

- 10 MR. TYLER: Ms. Cornich, could you pull up
- 11 Exhibit-1A, Page 6?
- 12 BY MR. TYLER:

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- Dr. Brookes, could you tell us what this is? 13
- 14 This is a Western blot from my laboratory, albeit quite
- 15 ugly one, but this is measuring the amount of a protein called
- ALK BH7. You see that down in the bottom right. A-L-K B-H 7. 16
- 17 We call it ALK BH 7. The date is in the top left corner.
- four blue marks that you see around the edge of the blot, those 18
- 19 mark the corners of the membrane. On the left, you see a
- 20 series of bands that have been marked with blue and red.
- 21 are the molecular weight markers which were transposed from the
- 22 membrane.
- 23 And then in this case, we are comparing two different
- 24 types of mice. So, if you read along the top, you can see that
- 25 every pair of lanes, one says WT, that stands for wild type.

1 That is just the normal laboratory mouse. And the other band

- 2 next to it says ALK BH 7 minus slash minus. That is a mouse in
- 3 | which that ALK BH 7 protein has been genetically deleted or
- 4 knocked out.
- 5 **Q.** Dr. Brookes, with respect to the molecular weight markers
- 6 which you mentioned a couple of times, what is the purpose of
- 7 that?
- 8 A. The purpose of that is to essentially tell us which band
- 9 to look for and to know that we are looking at the right
- 10 protein. So, it is fairly common, as you see here, this
- 11 Western blot was developed with an antibody against ALK BH 7.
- 12 But there are lots of bands. Not just a single band. So we
- 13 | need to know which band to look for and so reading across the
- 14 | lower red mark here, this is the 25-kilodalton molecular weight
- 15 marker. And if we read across, we can see that in particularly
- 16 the mito ALK BH 7 is a protein that resides inside
- 17 | mitochondria.
- So in this third set of lanes we see there is a prominent
- 19 band of 26 kilodalton in the wild type which is gone in the not
- 20 kept. And so that indicates the antibody is recognizing a
- 21 | protein at the correct molecular weight and that band is gone
- 22 | in the knock-out mouse. Therefore, that's the protein that we
- 23 | need to be looking at.
- 24 \mathbf{Q} . If you were to go ahead and publish this, how would you be
- 25 able to figure out what the size of the kilodaltons was at that

point?

- 2 A. So the relationship between the distance moved down the
- 3 gel in millimeters is a logarithmic function of the molecular
- 4 | weight, so we literally measure down the gel with a ruler and
- 5 | we plot a graph of distance versus the log of molecular weight,
- 6 and then we can read across and read down the gel and go to
- 7 that graph and know the molecular weight of protein.
- 8 Q. Now, Dr. Brookes, during the course of your work on this
- 9 case, did you develop a report to discuss some of the tools
- 10 | that you used to do your analysis?
- 11 **A.** Yes. There were three reports at the beginning that
- 12 described all of these various processes.
- MR. TYLER: And, Ms. Cornich, if you can bring up
- 14 | Exhibit-51?
- 15 **THE WITNESS:** Can I raise those -- how do I raise --
- 16 | thank you.
- 17 BY MR. TYLER:
- 18 **Q.** Dr. Brookes, could you tell us what this is?
- 19 **A.** So this is a walk through that describes some of the
- 20 techniques that are used in order to find the irregularities or
- 21 other problems in Western blot images.
- 22 **Q.** Are these some of the techniques and tools that you use in
- 23 your analysis in this case?
- 24 A. Correct. Yes.
- 25 MR. TYLER: If you can go to the next slide.

BY MR. TYLER:

- 2 Q. Dr. Brookes, if you could just walk us through this slide
- 3 show?

- 4 A. Okay. So, there are two images in this case. Image two
- 5 on the right is from the paper. Image one was obtained from
- 6 the data set. And so the first step is to bring both of these
- 7 | images in to Microsoft PowerPoint.
- Next slide, please. This is just a screen capture from
- 9 Microsoft PowerPoint. We're selecting the image on the right
- 10 and then we can bring up the format picture dialogue. And as
- 11 you see over on the right using the red arrow, there are a pair
- 12 of sliders. We can slide left and right to adjust the
- 13 | brightness and the contrast in order to enhance features in the
- 14 image.
- Next slide, please. This is an example of that. This is
- 16 | with the brightness turned down by 49 percent and the contrast
- 17 | increased by 78 percent.
- 18 Next slide, please. We can then crop out using the crop
- 19 | function in PowerPoint we can crop away the next.
- 20 Next slide, please. And, again, next slide.
- 21 This allows us to basically bring the figures next to each
- 22 other for a better comparison. So this is then the result of
- 23 | that. And, as you can see, it's highlighted in the red box.
- 24 There are a number of common features and the noise in the
- 25 background of these images which lead us to conclude that they

are, in fact, originate from the same image.

It's notable the red box is just shown as an example here,

but you can essentially pick any area in the entire background

of this image and all of the noise pattern is identical between

these two images.

A similar technique can be used in Adobe Photoshop if that is the preferred software. We can load the image in to Photoshop as it's shown mere.

Next slide, please. And we pull up the curves function from the menu. That brings up a dialogue box.

Next slide, please.

In the curves function, you will see a graph. The X axis of the graph has a scale from black to white all the way through gray in the middle. The Y axis of the graph has a similar scale, so the X axis of the graph is basically the input. It's saying, if you see a pixel of this density or this darkness, then apply an adjustment to it to give the output on the left.

So, in this case, there is no adjustment because the graph itself is a straight line, but the dot in the middle of the graph we can move up and down and left and right in order to create a complex profile that is akin to adjusting the brightness and contrast of the image.

If you go to the next slide, please.

So, this is a complex curve that has been applied and, as

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you see it, not only brings out differences in the background 1 of the image, this can also be used to enhance features within 2 the individual bands in the middle of the bands which 3 previously appeared pure black. 4 5 So this is a somewhat more useful feature than simply brightness and contrast. 6 7 Next slide, please. 8 Another example that can be used to pick out features in 9 blots is something called a gradient map. This is in Adobe 10 Photoshop again. 11 So this is under the same menu in Photoshop, the 12 adjustments menu, we click gradient map and that brings up a 13 dialogue box where we are faced with a number of different 14 color of choices. We can select any number of gradient maps in 15 order to enhance or bring down features of the image and, again, it should be known we are adjusting the entire image 16 17 here, not individual parts of the image. All of the image is being treated equally. 18 19 On the next slide, you can see three examples of that. 20 These are just three different color gradient maps that have 21 been applied. Some of them pick up different features than 22 other ones. And so these will be used then in a similar way to 23 compare background features, background noise, band, smears, 24 smudges in there. 25 Next slide, please.

If in the rare situation that we can't find an appropriate gradient map from that large list of possibilities, there is 2 3 the possibility here to go in to what is called the gradient editor where we can slide those color sliders left and right in

That is equivalent to essentially applying the curves function in order to find something that will pick up on certain features.

order to find an appropriate threshold or adjustment.

Next slide, please.

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So the last example here, this is an example of using something in Adobe Photoshop which is called a droplet.

Next slide, please.

So a droplet is essentially a macro or a plug in. set of predefined instructions telling Photoshop what to do and the droplets in this case are distributed by the Office of Research Integrity. This is the webpage where anybody can go download these droplets from ORI, plug them into Photoshop and it will perform this set of actions automatically with very little user input as a standardized way of doing this.

Next slide, please. So this is an example of that. is figure one from the grant application.

Next slide, please. This same figure was, in fact, also shown in a paper in Neurobiology of Aging. You can see by just looking at the band pattern that these images are similar, but what the goal of this analysis is to show that they actually

share a common digital heritage.

Next slide, please. So this is a PowerPoint file as well, so we can actually take the PowerPoint file which was used to prepare the grant for the paper image.

Now, in this case, Photoshop cannot recognize PowerPoint files. One is made by Microsoft and one is made by Adobe and they don't play nice. So we have to actually extract the image from the PowerPoint file in order to be able to analyze it in Photoshop. So that is done simply by right clicking and saving it.

The image that we are going to compare that to is this image here. This is an image that was provided to me during the investigation in which there is an area in the top right corner where a white box appears, and we have taken part of that image, highlighted it in red and cropped it and saved it, and that is the other image.

Next slide, please. So this is both images loaded in to Adobe Photoshop. Over on the right is a series of menus. This is called the actions menu. You can read the word actions in tiny tiny print at the top. These — this long list of things these are the various droplets. They come as a package from the ORI.

So the first step is we select the droplet that we want which is open at the moment, and down in the far bottom right corner you see there is a little play icon just like on your

music player, so we hit the play icon. 1 Next slide, please. And the first step is the macro runs. 2 It does some basic transformations to the image first. 3 makes a gray scale and then brings it back. And then applies a 4 5 gradient map to the image. So this is a red gradient map. These are preset within the ORI droplet. 6 7 On the next slide you can see we are applying a different 8 gradient map. Blue in this case or cyan to the other image. 9 Next slide, please. The next step is we actually bring 10 the images together. So in this case the image on the right 11 has been copied and pasted over into the image on the left. 12 That results in the two different images showing up in the 13 left-hand window as different layers. And the first thing we're asked to do is then apply a 14 15 curve so you see again a curve dialogue box, so we apply a curve to the first layer. 16 17 And the next slide. Then we apply a curve, the same The numbers in the curve boxes are the same so we apply 18 19 the same curve to the second layer. 20 Next slide, please. And then now that we have got both 21 images, both with the gradient map, both with curves applied,

images, both with the gradient map, both with curves applied, we can have them in the same window, and we have this double-layered image that we can use for comparison.

Next slide, please. The next step is then to apply what

is called a free transform to layer one. All that is is

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basically a dialogue that allows us to move and resize the image in order to line it up against the other one.

Next slide, please. So this is, for example, the image that's been resized. This is before the images have been aligned properly.

What you can see here is that anywhere there is red that is contribution for one of the images. Anywhere there is cyan that is contribution of the other images, but then anywhere within this white box in the middle, if there was any overlay between the band content that would show up as black.

And so on the next slide you can see that is essentially exactly what happens. When the two images, those — all that has happened between that previous slide and this slide is to just move one image over the other image. When they line up perfectly, all the red and the blue disappears, and we have a perfect alignment with black. And this indicates that these are, in fact, the same image. And we can blow that up and look at it on the next slide I believe.

One final step. If you want to save this, we merge the layers together and then on the next slide we use the save as dialogue box so we now have a single image. We save it as just a JPEG file.

On the next slide, please. Once we have saved it, this is then a blow up of the area of interest and so as you can see, there is basically no red and cyan here. It's all black.

1 These images align perfectly.

- 2 Q. Just one additional question on this, Dr. Brookes. With
- 3 respect to the black, is there -- did you use a different color
- 4 when you were doing your other reports?
- 5 A. Yes. Sometimes depending on, you know, which install of
- 6 Photoshop you are using, the presets of the gradient maps that
- 7 | are applied are slightly different, so I believe in some of the
- 8 | analyses the overlap appears in cyan or red, but the principle
- 9 is essentially the same.
- 10 Which color is which is described in the text of the
- 11 reports.
- 12 **Q.** And this slide show we just went through, is that
- 13 | something you prepared?
- 14 **A.** Yes.
- MR. TYLER: At this time the government would like to
- 16 move into evidence Exhibit-51.
- 17 | THE COURT: I thought it was already in. No?
- 18 MR. TYLER: This is one of the ones that was reserved
- 19 up.
- THE COURT: Okay.
- 21 MS. BEIDEL: Could we understand when it was
- 22 prepared?
- 23 THE COURT: Yeah. I'm actually not sure what this
- 24 is. I mean, is this --
- 25 MR. TYLER: This is just a --

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THE COURT: It's going to be used in the trial. 1 this just an example of how things work by using Dr. Brookes' 2 own data, Dr. Wang's data, somebody else's data? 3 Maybe you should ask more questions on that front. 4 5 MR. TYLER: Yes. BY MR. TYLER: 6 7 Dr. Brookes, when did you prepare this? 8 This was prepared last week. 9 And the images that we're looking at, whose images are 10 these? 11 These are images from the case that were provided to me, 12 so they're being used here purely on an example basis to 13 demonstrate the techniques. They happen to be very good 14 examples of images that overlap. 15 So it's not necessarily something you THE COURT: would use, Mr. Tyler, not necessarily something you would use 16 17 at trial, but just to illustrate the technique? Yes. And if we were to use it at trial, 18 MR. TYLER: it would be a demonstrative just to show how this works. 19 20 MS. BEIDEL: Your Honor, my understanding is that 21 this report corresponds to Dr. Brookes' previously produced 22 report 4.8. However, that 4.8 report does not use this droplet 23 method. 24 Our view is this has been manufactured very recently,

wasn't disclosed in expert discovery, and it's not appropriate

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for Dr. Brookes to be adding additional methods to bolster his
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    opinions at this time, so we object to the admission of this
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    document.
              THE COURT: What about that? Does this come from the
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    prior report or is it a new analysis?
              MR. TYLER:
                          It is a new analysis, but from our
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    perspective, Your Honor, it's just showing how the tool works.
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    It is not an opinion in and of itself.
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              THE COURT:
                          Were these particular droplets used in
10
    the analysis that you plan to use at trial?
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              MR. TYLER:
                          Yes.
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              THE COURT: In this exact same fashion?
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              MR. TYLER: Yes, Your Honor.
              MS. BEIDEL:
                           That's incorrect from what was disclosed
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    to us, Your Honor. There's either a disclosure issue or that
16
    is not a correct representation.
                          The example -- the techniques -- and we
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              MR. TYLER:
    can walk through this in some other examples, Your Honor, that
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    we have. The technique is something that will be used at
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    trial. This specific PowerPoint presentation was just put
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    together as an example of how --
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              THE COURT:
                          Why weren't we just using the one that
23
    you are going to use at trial? It seems to me that would be
24
    easier.
25
              MR. TYLER:
                          The one we are going to use at trial --
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THE COURT: Are they different in some way?

2 MR. TYLER: Essentially it is just the last slide

3 here. So, for Your Honor's benefit, we are trying to show what

the steps are to get to that last slide to show how that

5 actually came to be.

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THE COURT: Okay. Well, I mean, I think for purposes of this hearing we are trying to understand the techniques and so forth, I'm going to allow it. But, again, it does not mean we are going to use it at trial and I think we should have cross-examination on sort of how this compares or does not compare to what was actually done, and I generally agree with the principle that if it's not something that was disclosed previously, you are going to have a problem getting it in at trial so.

But for purposes of just understanding the technique -- again, it's not clear to me this was the same technique used, but we will find out after cross-examination.

MR. TYLER: Thank you, Your Honor.

THE COURT: Go ahead.

- 21 **Q.** Now, Dr. Brookes, turning to the source of the materials
- 22 that you used for your actual analysis, what were the sources
- 23 of materials that you used?
- 24 | A. In the case of grants and source images, they were sent to
- 25 me by Agent Jeffrey Weeks of the FBI.

Q. And what were those?

- 2 A. They were PDFs of grants, as well as JPEG images. In
- 3 addition, in some cases, there were papers. I was able to
- 4 obtain those papers myself because they are freely available
- 5 and public through my university library subscription.
- 6 Q. In addition by e-mail, is there other ways you obtained
- 7 information from Special Agent Weeks?
- 8 A. Yes. I was sent a pair of DVDs containing one DVD
- 9 contained approximately 2,300 JPEG images and another DVD
- 10 | contained approximately 2,900 PowerPoint files.
- 11 **Q.** I apologize if you already said this. But did you also
- 12 | receive the grant applications?
- 13 | A. Yes. As I mentioned, the grant applications were in PDF
- 14 format sent by Agent Weeks.
- 15 Q. Turning to before -- I guess last thing before we get to
- 16 the analysis, which is Dr. Brookes, did you do a report which
- 17 described your general method of how you plan to approach this
- 18 | analysis?
- 19 | A. Correct. Yes. The general methods that we applied were,
- 20 | first of all, we -- sorry. I keep saying we. I. I had a PDF
- 21 of the grant with a figure in the grant that was the figure in
- 22 question, as well as image that was known as a white box image.
- 23 | It will become apparent what we are talking about in a second.
- So the first step was to try to work backwards from the
- 25 grant to see if any -- if the white box image was, in fact, the

source for the grant image, so that was -- that was termed a reverse analysis. So the objective there is working backwards to try to trace the chain of provenance for the data in the grant.

In addition to that, I performed a forward analysis in which I was presented with a related raw image, a raw Western blot with no annotation or white box on it. Working forward hypothesizing, can we create the image in the white box or the final published grant figure from the raw image, so hypothesizing whether that can be done.

- Q. Generally speaking, how does that approach align with the scientific process, in your view?
- A. Essentially -- I mean, it aligns on a couple of different manners. One is when we're performing a reverse analysis, we are essentially trying to establish as scientists, when performing these kind of experiments, what is the chain of custody or the chain of provenance of the data. It's an essential part of the scientific process. The data should be traceable to the raw data from which it came.

And then in the forwards case what I'm essentially doing is starting with the hypothesis. The hypothesis is that this image can be manipulated in a fair and standardized manner using approved tools in the field, such as adjusting the brightness and the contrast evenly across the entire image, and that will bring about the bands or the pattern of bands or the

final data that is in the white box or the final published image.

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Q. Dr. Brookes, in that -- not in that report, but in other reports did you consider potential alternatives, explanations for your analysis?

A lot of alternative explanations were explored. One in particular which I don't think is possible. Western blot is run, quite often if the result turns out messy, maybe there is a fingerprint in the middle of the film or a band didn't come out properly, it can be fairly common to use a process which is called stripping and reprobing. So we take the membrane, wash away all the antibody and go back with a fresh match of antibody and redevelop a completely brand new piece of film. That's called stripping and reprobing. possible, but what that does is it results in a completely different background image because when you strip and reprobe, you are taking the membrane out of the cassette, bits of dust, bits of hair, whatever, get on it. It's going to get scratched, washed around in the dish. When we put it back into the cassette, it's going to be in brand new wallet binder which

So, when a second piece of film is developed from that membrane, there is absolutely no chance that the exact same pattern of noise can be reproduced on that second piece of film.

has got different creases and things on it.

So stripping and reprobing was one specific alternative that we ruled out as a possibility.

- MR. TYLER: Now, turning to the analysis.
- 4 Ms. Cornich, could you unplug Exhibit-39, Page 60?
- 5 BY MR. TYLER:
- 6 Q. And, Dr. Brookes, could you tell us what we're looking at
- 7 here?

- 8 A. This is a page from an N.I.H. grant and figure one of the grant shows an isoelectric focusing gel.
- MR. TYLER: Ms. Cornich, if you could also bring up

 11 Exhibit-37, Page 7.
- 12 **THE WITNESS:** At the top of the page on the left you see the same data published in a paper in the Journal of
- 14 Neurobiology of Aging. These are the same data as shown in the
- 15 grant.
- 16 BY MR. TYLER:
- 17 | Q. Dr. Brookes, could you explain just what the -- basically
- 18 | what the information being conveyed here is?
- 19 | A. Yes. So this is an isoelectric focusing gel. Along the
- 20 | left-hand edge, you can see there is a scale. That is a pH
- 21 | scale. So 9.5 means the isoelectric point at which the protein
- 22 has no charge would be a pH of 9.5 and the same down at the
- 23 | bottom is 3.5. So, typically, when these experiments are done,
- 24 | they are done on a pH gradient of three to ten. Three to ten
- 25 is the maximum range here.

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In the first three lanes of the Western blot, the first three bands that you see, those are from controlled samples. The second set of three bands are from Alzheimer's Disease samples. Along the bottom, you can see PTI 125 or animalia. Where there is a plus below the third lane, that indicates that's a sample which has been treated with the drug PTI 125. What you can see by comparing the first lane and the fourth lane is that in a controlled sample the isoelectric point of the protein is 5.9. It roughly lines up with six on the left-hand scale. In the Alzheimer's sample, it has shifted down to about 5.2, 5.3. This is used to make the claim that Alzheimer's results in a confirmational shift of the protein. The reason isoelectric focusing is done in this case is because, as I mentioned, all proteins have a native negative charge with positive and negative charges, but when proteins fold differently, that can cause, for example, one area of a protein to be covered up and that can obscure positive or negative charges, and that can change the isoelectric point. So we can see in the fourth lane across the band has shifted down. If we then turn our attention to the sixth lane, what appears to happen in the sixth lane --THE COURT: Can you just describe what you mean by "lane"? Can you point or circle? You can mark it. THE WITNESS: A lane is a vertical designation.

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So maybe you can just point to the ones THE COURT: 1 2 you are referring to.

So this will be lane number one. THE WITNESS: will be lane number three versus four. This is lane number six.

> THE COURT: Thank you.

THE WITNESS: And so comparing the middle two lanes, three versus four, you can see Alzheimer's Disease introduces a shift in the isoelectric point of the protein. It's moving down the gel further. And then in the very last lane PTI 125 over on the far right appears to reverse some of that shift. It brings some of the protein back up the gel to make it more similar to what we see in the control condition.

BY MR. TYLER: 14

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- 15 So, in layman's terms, what does that mean?
- 16 So that is -- that is essentially saying that the drug 17 appears to reverse the conformational change that is happening in the protein in Alzheimer's Disease. 18
- 19 And then, also, Dr. Brookes, can you explain the 20 relationship to the bar charts in the middle?
- 21 Yes. So the bar chart in the middle is essentially a 22 quantitation of the amount of each of the two different 23 isoelectric points.
- 24 So the white bars would be the density of a band with an 25 isoelectric point of 5.9. The black bar is the density of a

band with an isoelectric point of 5.3. And as you see here in the control, just as in the control here, there are all of the protein appears to be in the 5.9 state as indicated by the fact

that there are tall white bars with no black bars.

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However, again, in Alzheimer's in the fourth lane there appears to be a shift where now most or all of the protein is in the 5.3 state, so now the white bar has gone down. The black bar has come up. And then over in the far right you can see that there is a mixed population. There are both white and black bars as if the -- some of the protein that was in the lower state has been shifted to the higher state by the drug.

This graph here is a quantitative graph of the density of these bands and you can see in the figure legend down here it states N equals six. N in biological sciences when we are referring to this in science, the N is the number of replicates.

So the figure legend is stating that these data are the average of six independent times that this experiment was performed.

- Q. Dr. Brookes, could you explain what the little like T looking bars on the top of the bars, what that indicates?
- 22 **A.** Yes. So when we -- when we perform these types of experiments, you never get the same answer twice. Right.
- 24 There is a joke among researchers, if every experiment worked

25 | the first time, it wouldn't be called research.

And so the quantification of the bar, the density, the average is given by the height of the bar. The little T thing

3 above is called an error bar, and that is an expression of the

4 range or the variance of the result.

5 So a very small or tight error bar, as you see here,

6 indicates an experiment that gives you the same result every

7 time. The result is very reproducible and tight. A bigger

8 error bar would indicate more variance in the result and

9 because experiments are varied, it's often necessary to perform

them multiple times such as six times in order to see a

11 significant difference.

12 Q. Thank you, Dr. Brookes. This figure which we noted here,

13 did you see this reproduced across more than one grant

14 application?

10

15 A. Correct. Yes. I believe four different grants.

MR. TYLER: Now, Ms. Cornich, if you could pull up

17 Exhibit-10A?

18 **THE WITNESS:** Can we delete the -- thank you.

19 BY MR. TYLER:

20 **Q.** Dr. Brookes, could you explain what this is here?

21 | A. So this is a report that I authored regarding figure one

from the N.I.H. grant proposal.

23 MR. TYLER: And, Ms. Cornich, we can go to the next

24 slide.

22

Q. And Dr. Brookes, what is -- what do we see here?

2 A. This is the figure from the grant from the paper. They're

the same figure.

MR. TYLER: And Ms. Cornich, if we could go to the

next slide?

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- Q. Dr. Brookes, could you tell us what we see here and what
- 8 you did here?
- 9 A. Okay. So on the top right where it says C, final
- 10 published, that is the image from the grant. Over on the left
- 11 there is an image that I call the white box image. You will
- 12 see from the title of this JPEG file, C is controlled. AD is
- 13 | Alzheimer's Disease. FLNA is Filamin A conformation. IEF,
- 14 this is an isoelectric focusing gel-11. Numerous of the white
- 15 | box images that I encountered in this analysis had the number
- 16 | 11 appended to the file name.
- 17 The whole image, as you see it, is essentially a scan of a
- 18 piece of film. It's capturing a piece of film like this. The
- 19 | top right corner of this image appears to contain white boxes
- 20 of unknown problems. If we take a section of that white box
- 21 | image as shown in red and we enlarge it and we subject it to a
- 22 | vertical stretch and place it below the final image, so here on
- 23 the right, you can see from the areas highlighted in blue that
- 24 the pattern of bands matches exactly between the two images.
- In addition, the series of smudges above and below the

bands and the background noise is also a match.

2 And so from this, I conclude that the white box image on

3 the left is the digital source for the final published image.

MR. TYLER: Ms. Cornich, if we can go to the next

5 slide.

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6 **THE WITNESS:** This is essentially showing the ORI

7 droplet overlay.

So, as was mentioned previously, when they overlay, it was in black. On this particular slide red happens to be the color

where overlay is indicated. As I mentioned, it just depends on

11 what default gradient match you have installed in Photoshop

12 determines what those eventual colors are, but the end result

13 is the same which is that these two images overlap including

not only the bands, but also the noise above and below the

15 bands.

- 17 | Q. Dr. Brookes, I probably should have picked up on this
- 18 | earlier, but is this example actually the one we went through
- 19 in that Exhibit-51 earlier?
- 20 **A.** Correct. Correct. This is the same example.
- 21 | MR. TYLER: Ms. Cornich, if we can go to the next
- 22 page.
- 23 BY MR. TYLER:
- 24 | Q. Dr. Brookes, can you tell us what kind of comparison you
- 25 are doing here?

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So on the right is the white box image that you saw Α. previously. You'll note the file name with the minus 11 up at the very top right. On the left is a related image from this family of images. This is the raw image. A number of features about this document. First of all, you will see in the file name it's missing the number 11. But, otherwise, the file name is identical. In addition, as shown, this image appears to have a lot of commonalities with the white box image in the areas outside of the white boxes, so essentially you can look at anywhere outside the white boxes on the left image and you will find the corresponding area in the right image. images are essentially identical, all apart from the area in the white box. In terms of where this image came from, the image on the left as shown in red, it appears to have rotational symmetry, as well as vertical symmetry as shown in blue. So what does that mean? So, in theory, we could fit eight Western blots on a single piece of film. However, that's a lot of work, so quite often we do two Western blots. And so this is thought to be an image with two Western blots and so, for example, you will expose the film for ten seconds, turn the gel around, expose it for 30 seconds, turn the gel around, expose it for a minute, flip it around and expose it for two minutes. So you can capture four different exposures of the same pair of membranes on a single piece of film. And as you can

see, that results in slight differences in darkness for the different exposures.

So it's a bit of a mental loop, but if you look over in the top left corner, there is a membrane and then in the bottom right corner is the same membrane. Those are the same membrane, just two different exposures, and you will see one of them is — the one in the top left is slightly darker than the one in the top right.

So this is essentially four different exposures of the same two membranes. So because of all the commonalities between the raw image and the white box image in terms of everything outside the area of the white box, it is concluded that this raw image is the source digitally for the white box image.

- Q. Would you expect to see any markers on the raw image?
- A. Yes. That is an unusual features of this image. In the final published image, there are pH markers for isoelectric point. Those are used to calibrate and essentially say something about the conformation and the isoelectric point.

However, neither the white box nor the raw image appear to have any pH markers. There are no annotations handwritten on the membrane or the piece of film, so it's difficult to see where those markers in the final figure originated from.

- Q. Is that something you have seen, in your experience?
- 25 | A. No. That's highly unusual. That is not common standard

1 | scientific practice. You annotated your gel so you have, as

- 2 was mentioned, a chain of provenance for the scientific data.
- 3 MR. TYLER: Ms. Cornich, if we can go to the next
- 4 page.
- 5 BY MR. TYLER:
- 6 Q. Dr. Brookes, can you tell us what we are looking at on
- 7 this page?
- 8 A. So this is now honing in on the top right corner of each
- 9 of those images. The image on the right is the white box. The
- 10 | image on the left is the raw. And as you can see from the area
- 11 | highlighted in red, the noise pattern is reproduced from left
- 12 to right. As you see in --
- 13 **Q.** Sorry.
- 14 **A.** As you see in -- from left to right between the raw image
- 15 and the white box image, the area in red is essentially
- 16 | identical. The noise pattern, the fingerprint of noise from
- 17 the piece of film is transferred perfectly to the white box
- 18 | image. However, if you look at the position in the white box
- 19 | image where the bands appear which is in blue, there are no
- 20 corresponding bands in the left-hand image.
- MR. TYLER: Now, Ms. Cornich, if we can go to the
- 22 next slide.
- 23 BY MR. TYLER:
- 24 | Q. Dr. Brookes, if you can tell us what we are looking at,
- 25 | what analysis you did here?

A. So this is the forward analysis that was mentioned in this experiment, and essentially attempting to manipulate the raw image using fair and standardized methods that are accepted in the field. That is adjusting the brightness and contrast of the whole image evenly. An entire series of brightness and contrast adjustment were made. This just shows one example.

We don't have room to go through every one percent step of

We don't have room to go through every one percent step of both.

So what I was hypothesizing here is that perhaps there are bands hidden in the raw image and that I might be able to adjust the raw image in such a way as to make the bands in the white box image appear.

What is apparent from what you see in red is that, yes, indeed, some of the noise surrounding the bands, the dot in the middle, the smudge over in the far right, some of the noise does, indeed, appear to transpose from the white -- from the raw to the white box image. However, no adjustment that I was -- that I performed was capable of making the bands in the white box image come in to appearance.

- Q. And what does that show about the ability to recreate the bands in the white box image from the raw image?
- A. So the fact that the noise transposes proves the digital provenance here. The white box image is a digital child of the raw image. However, the bands don't have prominence in the raw image.

- 1 Q. The image on the left, is that an appropriate
- 2 | representation then of the raw image?
- 3 **A.** That is an adjustment of the raw image, so anybody who
- 4 opened this in PowerPoint would be able to take that raw image
- 5 and adjust the brightness and contrast back to zero and arrive
- 6 back at the raw image so, yes, the image on the left does
- 7 | represent the raw data, albeit with the brightness and the
- 8 | contrast adjusted.
- 9 \mathbf{Q} . And in your view and experience, is that publishable data?
- 10 **A.** The data on the left, in my opinion, is not publishable
- 11 | because it does not appear to contain usable bands. And, in
- 12 particular, if we go -- if we count across four lanes. If we
- 13 go one, two, three, four or five. If we look in lanes four and
- 14 | five, there do not appear to be any bands at all. There are
- 15 some smears and smudges. However, in lanes four and five of
- 16 the final published image, there are prominent sharp bands and
- 17 | it's impossible to tell where they came from.
- 18 MR. TYLER: Ms. Cornich, if we can go to the next
- 19 slide, please.
- 20 BY MR. TYLER:
- 21 **Q.** Dr. Brookes, can you tell us what we see here?
- 22 **A.** This is just essentially the same as you saw on the
- 23 | previous slide, just using gradient maps to illustrate the same
- 24 point. So, for example, if we -- the red boxes on the previous
- 25 | slide, they appear black in this example.

- So, if we look at the right hand red box in each case, you
- 2 can see the sort of little pattern of noise that looks like New
- 3 Zealand with some islands off the shore, that is a unique
- 4 imprint of this image, that noise being the same is a traceable
- 5 fingerprint between these two images that indicates that they
- 6 are digital relatives. So this is essentially just another way
- 7 of showing the same point.
- 8 Q. Did you consider other possible explanations for what was
- 9 going on here?
- 10 A. Yes. Other things such as stripping and reprobing. We
- 11 also considered the possibility that there may be additional
- 12 | images that exist in between the raw image and the white box
- 13 | image. However, that does not change the fact that the raw
- 14 | image and the white box image are digitally related.
- So, regardless of how many steps were involved in between
- 16 the raw image and the white box image, it still would not
- 17 explain where the bands came from in the light box image
- 18 because they were not there in the raw image.
- 19 MR. TYLER: Ms. Cornich, will you go to the next
- 20 slide.
- 21 BY MR. TYLER:
- 22 **Q.** Now, Dr. Brookes, could you tell us what analysis you are
- 23 | doing here on this slide?
- 24 A. Yes. So I apologize. This is going to get a bit
- 25 technical. So over on the right, we have the published image.

That is the isoelectric focusing. The pH scale runs from 3.5 to 9.5. As I mentioned, when you run an isoelectric focusing gel, the gel occupies the entire -- the pH gradient occupies the entire height of the gel.

So, when you do an IEF gel, the top of the gel is ten.

The bottom of the gel is three. That is an incontrovertible scale. You can't do anything about that. So, remember, the bottom of the gel is three.

If we go over to the white box image, you can see we have taken the whole of the gel, the full height of the gel in the blue box. And if I subject that blue box gel to a vertical stretching, in order to make the bands line up, so the bands here line up, you can see, in fact, that the blue gel scaled, if it were real, would cover much more than three to ten. In fact, it looks like it would go all the way down to pH zero which is physically impossible. It may even go to pH minus one. There is no such thing as a negative pH.

So, based on this, I conclude that the stated pH range here is physically impossible. Three to ten cannot had been the original values on this full scale height of this blog shown in the white box image.

So this particular pH scale is not only do we not know where it came from, but it's also impossible given the relative scaling of these things and the vertical stretching that has occurred.

- Q. Where would you expect to see that or that scale, that pH scale on the white box image?
- 3 A. I would expect to see some markers on the white box image
- 4 or on the raw image if they show up in the final published
- 5 image.
- 6 Q. And, in your experience, is there -- are there other ways
- 7 | that you can transfer that information?
- 8 A. If you wanted to, you could type that or you could try to
- 9 | note it down in a book or have the film stuck to a piece of
- 10 paper, but then you -- you know, any time you are not keeping
- 11 | the data together all in one place, that risks mistakes, that
- 12 risks sloppy lab practices.
- So, for example, here everything is in one piece of paper.
- 14 This piece of film contains the data, contains the molecular
- 15 | weight markers. If that gets run over by a car, the data is
- 16 still there. We are not going to lose any of that data.
- 17 Whereas, if it was kept in a separate place, if the molecular
- 18 | weight markers or the pH markers were in a different place, in
- 19 different file, in a different folder that creates the
- 20 likelihood of mistakes further down the road.
- 21 So the scientifically correct way to do this is to keep
- 22 all of the information in one place together.
- 23 | MR. TYLER: Ms. Cornich, now if we could turn to
- 24 Exhibit-10B.
- 25 BY MR. TYLER:

- Q. Dr. Brookes, can you tell us what this document is?
- 2 **A.** So this is a PowerPoint file that was provided by Agent
- 3 | Weeks. On the first page of the PowerPoint file is what
- 4 appears to be the figure for the paper. It's quite common to
- 5 take a Western blot image, import it in to PowerPoint and then
- 6 apply the various annotations as you see here. The scale, the
- 7 labels in a software such as Microsoft PowerPoint.
- 8 MR. TYLER: If we can go to the second slide.
- 9 BY MR. TYLER:

- 10 **Q.** Dr. Brookes, did you discover anything about this slide?
- 11 A. So this slide contains the graph, as you see from the
- 12 paper and the grant. These appear to be the same data.
- 13 Everything about it is the same including, for example, the
- 14 | minute positioning of the stars used to indicate significance
- 15 | so that this does appear to be the figure. If we right click
- 16 on this figure, we can go to chart object, and we can select
- 17 open and that will actually export a marker off the Excel file.
- 18 And so there are several ways to get a graph from
- 19 Microsoft Excel in to PowerPoint. However, when that is done
- 20 directly by just dragging it into PowerPoint, all of the Excel
- 21 | file and the data used to plot the graph comes along for the
- 22 ride. So you can see in this Excel file which was just
- 23 | generated de novo right here in this courtroom from the
- 24 | PowerPoint, if we go to sheet one, this is, if we scroll up,
- 25 this is the data that was used to generate that chart that

appears in the grant and the paper.

- 2 MR. TYLER: Ms. Cornich, if we can now pull up
- 3 Exhibit-10C.

- 4 BY MR. TYLER:
- 5 Q. Dr. Brookes, can you tell us what this is and what its
- 6 relationship is to the Excel sheet we were just looking at?
- 7 **A.** Yes. So, over on the left you will see PI 5.9 and 5.2 or
- 8 maybe it's a 5.3. It is not clear. There are two different
- 9 values there. The area in yellow --
- 10 **Q.** I'm sorry, Dr. Brookes, to interrupt you. What is this in
- 11 | comparison to the last spreadsheet and what is it says --
- 12 **THE COURT:** Can I ask, is this a slide that's in
- 13 Exhibit-10B, or is it something else?
- 14 MR. TYLER: It is -- Your Honor, this 10C is an
- 15 annotated version of the one that we just extracted.
- 16 **THE COURT:** I guess I'm not seeing a Bates number or
- 17 | anything like that, so I don't even know if this is part of
- 18 | what you just offered as 10B.
- 19 MR. TYLER: So 10B is a PowerPoint with the embedded
- 20 Excel sheet and 10C is that same Excel sheet annotated by Dr.
- 21 Brookes.
- THE COURT: So, when you say it was embedded, it's
- 23 electronically embedded, but the physical document, in theory,
- 24 is a piece of paper, so this is some sort of native file?
- MR. TYLER: Correct.

1 THE COURT: Okay. I mean, I think just so you know

2 where we are going to have to have this in a format where we

know what we are looking at. This may or may not be easily

4 replicable in the record somehow in the future, so you are

5 going to need to have screens of this, not just pulling things

up off your computer which can be changed.

MS. BEIDEL: Your Honor, in addition to that concern, I haven't heard a tie in to Western blotting methods at all and in the interest of time today, I'm not sure -- I feel like we are down a rabbit trail at this point talking about pH data.

THE COURT: I mean, there are things in the expert reports about this. There wasn't a whole lot of detail in the report, but I think this issue came up, I think. So why don't we go a little further and see where it goes.

MR. TYLER: Yes, Your Honor.

BY MR. TYLER:

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- Q. Dr. Brookes, could you just walk us through this annotated version of that spreadsheet?
- A. Yes. So over on the left in Column A we have the labels
 of the different isoelectric points. 5.9 and 5.2. The numbers
 over to the right are densitometry values. So these are values
 purporting to represent the density of the bands in each of the
 samples. There appear to be six sets of data so, for example,
 rows eight and nine those are one set of data. Row 12 and 13,
 that is a second set of data. So there are six sets of data.

These are highlighted in yellow. I should note that is not my highlight. The spreadsheet came like this, so this set of data

highlighted in yellow is what was used in the presentation.

As you will see in cell number C9, there is a five. That is not particularly unusual. However, a cross in cell number E9 and G9, and then down in number 13, you will see there are a number of other fives, and then further down there are tens.

with this frequency in stochastically acquired biological data. There does not appear to be an explanation for this. The simple one would be, oh, maybe this is a rounding error. Maybe everything gets rounded to the nearest five or zero.

It is very unusual that the number five and ten appears

However, the other numbers in the spreadsheet do not all end in five and zero, so that does not appear to be a reasonable explanation for why.

Secondly, for reasons that I really don't understand, there is a set of six independent experiments in the central yellow part of the sheet. If we could scroll up please to row eight and nine, you will see the values — the first value 1542. Reading across 1627. If we scroll down now, for some unexplained reason that same first set of data appears three more times in the spreadsheet.

Typically, a scientist, when we are trying to keep track of things, it's bad enough trying to keep track of things, but having three extra versions of one of the independent

experiments inexplicably included in the sheet for no apparent reason, that appears to be, again, just not following standard

Scrolling down further to the third, what has happened here in red and the numbers below in red is the size of those little T arrow bars is being calculated. So, if you click on some of D47, what's being calculated here is the average density of the six samples. And then in the cell below in D48, we are calculating the size of that arrow bar.

Now, the typical mathematical formula that is used to calculate that and the formula is written in the formula bar in Excel right here — the typical formula is to take the standard deviation of the numbers and divide it by the square root of the number of samples, so if the —

MS. BEIDEL: Objection.

practices. Very very unusual.

THE COURT: What's the objection?

MS. BEIDEL: There's no methodology for this type of analysis in the report. He has a methodology that is a three-step analysis of Western blot images. Now he's telling us about typical scientific methods for all kinds of things.

THE COURT: I agree these are issues, and I think for purposes of this hearing only, I think we should handle that through cross-examination first. And then if you want to try to exclude something at that point, we can. Obviously, we are handling this differently than we would at trial, but let's

handle it that way.

2 MS. BEIDEL: Understood, Your Honor. Thank you.

THE COURT: Thank you.

BY MR. TYLER:

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- Q. Please proceed.
- So, the typical formula -- in fact, the formula for the 6 7 calculation of the standard error of the mean is to take the 8 standard deviation and divide it by the square root of N, so 9 this would be the square root of six. For an inexplicable 10 reason here, the error bar has been calculated when multiplying 11 the standard deviation by five. Further over, on comment 12 number four, it's very similar and related. Again, instead of 13 dividing the error by a number, they multiplied the error by a 14 number to calculate the error bar. That is just not the way 15 that errors are calculated.

The wrong formula has been used to calculate errors. However, what is notable is that in every case the result is that the error bars appear larger than would be typical.

So I have calculated, again, one of the errors here in cell number Q48 using the correct formula. And what you see by comparison with the number above, the number above is 70 percent, plus or minus two percent. Two percent is the correct error. This is how it should had been calculated in the spreadsheet. Instead, the error that is reported in the graph is 9.7 percent over on the left in cell number 048.

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Generally speaking, biological processes are noisy.
    time we transfer liquid, we are patting. Everything is
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    calibrated, but there is noise. I would say on a good day it
    is very good to get anything less than a ten percent error on a
    biological experiment. A two percent error, on the other hand,
    would attract scrutiny from reviewers as being too small.
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 7
    two percent error would raise eyebrows --
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              MS. BEIDEL:
                           Objection. Speculation.
 9
              THE WITNESS: -- in the field.
                          So, again, we will handle this on
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    cross-examination I think to start with. But can I just --
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    again, Mr. Tyler, it's kind of confusing. I thought that what
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    we had here was a PowerPoint that was perhaps created by
    Dr. Wang with a spreadsheet. Now I'm looking at a screen where
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    there is all types of annotations by Dr. Brookes.
         So what is this? This isn't evidence, is it?
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              MR. TYLER:
                          No.
                               This is demonstrative to --
              THE COURT:
                          We need to have numbers and understand
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    exactly what we are looking at here because if this happened in
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    front of the jury, I would be probably excluding it entirely
    because you purported it to be actual evidence and now it's
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    just work product by the expert which are two totally different
23
    things.
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              MR. TYLER: Understood, Your Honor.
              THE COURT:
                          Can I just understand, is this
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spreadsheet something that supposedly was in the raw
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    information obtained from Dr. Wang?
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              MR. TYLER: Yes, Your Honor.
              THE COURT: And, then, these annotations were added
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    later for some unknown reason?
              MR. TYLER:
                          The annotations were added later as
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    analysis, Your Honor.
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              THE COURT:
                          Then why didn't you say that when you
 9
    offered it? You said this was the data.
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              MR. TYLER:
                          I'm sorry, Your Honor. When we switched
    from 10B to 10C, I included --
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              THE COURT:
                         When we switched to 10C, just this
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    spreadsheet or the PowerPoint?
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                         10C is the spreadsheet annotated by Dr.
              MR. TYLER:
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             10B is the spreadsheet that is derivative from
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    Dr. Wang's PowerPoint.
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              THE COURT: How are they different? Just the
    annotations?
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              MR. TYLER: Just the annotations and this calculation
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    that is PSB5 he's pointing to.
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              MS. BEIDEL: Again, Your Honor, I believe this
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    annotated version to have only been produced in connection with
23
    this hearing or just this week produced so, again, we have a
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    disclosure problem with respect to this, unless it's --
25
              THE COURT: Has the defense seen this before?
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It's not Bates labeled in the --MS. BEIDEL: 1 This has been disclosed before, Your 2 MR. TYLER: 3 Honor. How recently, Your Honor? 4 MS. BEIDEL: 5 MR. TYLER: This was disclosed as part of the original production. 6 7 The spreadsheet? THE COURT: 8 MR. TYLER: Yes. 9 THE COURT: With all the annotations, the bizarrely 10 comments and the like? 11 MR. TYLER: Yes, Your Honor. 12 THE COURT: Okay. 13 With respect to Dr. Brookes' material, MS. BEIDEL: 14 something we have been asking the government to do for months 15 now is to give us an organized version of an understanding of what Dr. Brookes' report is. There are all these materials 16 17 strewn throughout the two million pages of discovery materials that we can't possibly find everything. We have a right to 18 19 have an understanding of what the government's conception of 20 the report is. 21 THE COURT: We can certainly have this discussion 22 later without taking up Dr. Brookes' time. So why don't we 23 finish with the testimony? We will certainly have that 24 discussion. 25 MS. BEIDEL: Thank you.

MR. TYLER: Thank you, Your Honor.

BY MR. TYLER: 2

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- Dr. Brookes, if you could just do the last two comments here?
- So, scrolling over to the left, and scrolling up a bit and over to the left, essentially what was done after the data were 6 7 calculated, so in the upper set of red and blue bars of the 8 calculations of the average data from the six experiments.

Now, sometimes it's necessary to move those cells around in order to then plot the data in a graph. And so the numbers in red in the top set of bars were copied and pasted or moved somehow to give rise to the set of numbers in the red set of bars. And the data in this lower set of bars here in red and blue, that is what was used to plot the graph that you see in the grant.

What's unusual here is that while all the numbers in red and blue appear to track properly, so, for example, 0.994 in cell number D47 becomes 99.4 percent in cell number D56. everything in red and blue tracks properly, so there is a process by which the data that was calculated has been transposed into a format that makes it easier to select for plotting the graph.

However, the numbers in orange have not transposed and, in fact, for example, if we look at cell number M51, that is a 1.6 percent error. If we now look at cell I60, that is a

four percent error. That number four and the three next to it 1

and the 99.7 above it in orange, those numbers appear to have 2

- 3 just come from nowhere. They do not have any provenance
- elsewhere, unlike all of the blue and red numbers all of which 4
- 5 can be traced to the upper set of blue and red numbers which
- can, in turn, be traced to the rest of the data file. 6
- 7 So the numbers here that were used to generate those T
- 8 bars, those error bars in the final graph, these are the
- 9 numbers and they don't have any provenance anywhere else in
- 10 this spreadsheet. That three and four appear to me to have
- 11 been fabricated.
- 12 Dr. Brookes, would you expect to see a formula in that bar
- on the top, is that related to what your conclusion there? 13
- 14 Yes. I would expect if that cell is related to
- 15 another cell I would expect to see a formula telling us how
- 16 they are related.
- 17 MR. TYLER: Ms. Cornich, if we can go to the left all
- 18 the way and cover the last.
- THE WITNESS: So the last point turns to the 19
- 20 feasibility of some of these numbers. When we express the
- 21 average plus or minus an error bar, that error bar is
- 22 essentially expressing the range of possible values of the data
- 23 that contributed to that average.
- 24 And so, for example, here if we look in cell number D57,
- 25 we have a value of 99.4. The range is giving in D60 which is

So the way we express this scientifically is 99.4, plus

- 2 or minus 2.5. The fact of the -- if you take 99.4 and you add
- 3 2.5, you get a number that is bigger than a hundred. That
- 4 means that some of the numbers that contributed to that 99.4
- 5 average must have been bigger than a hundred percent. And that
- 6 is physiologically impossible. You can't have more than a
- 7 hundred percent of a protein in a particular confirmation.
- So, again, the error bars relative to the actual numbers are physically impossible.
- 10 Q. Thank you, Dr. Brookes.
- 11 MR. TYLER: Just to close a loop here, Ms. Cornich,
- 12 can you bring up Exhibit-10, Page 4.
- 13 BY MR. TYLER:

2.5.

- 14 Q. Dr. Brookes, could you just tell us what we're looking at
- 15 here?
- 16 **A.** So this is the text description in the report describing
- 17 | the PowerPoint file, where the numbers came from, the
- 18 extraction of -- so this is describing in text the process by
- 19 which the PowerPoint file was extracted and then a summary of
- 20 the annotations in the XL file that were just discussed.
- 21 Q. So that's a narrative of what we --
- 22 **A.** That's a narrative of what we just discussed, yes.
- 23 | THE COURT: Give me the number on this again, please.
- 24 | MR. TYLER: It's Exhibit-10, Page 4. If we go to the
- 25 | bottom, I can give you the Bates number. The last six of the

Bates are 352197.

2 **THE COURT:** So this is the expert report on 4.7?

3 MR. TYLER: That's correct, Your Honor.

4 THE COURT: That we already have. Thank you.

BY MR. TYLER:

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- Q. Dr. Brookes, taking on the analysis together with respect to figure one, what did you conclude with respect to the
- 8 research record here?
- A. So I conclude on the basis of this analysis that the bands that appear in the final published image and in the white box image find absolutely no provenance in the raw data. In addition, the graph the error bars in the graph in particular, appear to have been fabricated and, therefore, I concluded the result as it is presented in the published grant
- MR. TYLER: Your Honor, this is a little bit of a stopping point. I don't know if the court reporter needs a five-minute break. We are happy to keep going.

and paper does not represent the scientific record.

- THE COURT: How much longer do you have with this witness?
- MR. TYLER: As we have it set up, about 45 minutes.
- THE COURT: Why don't we take a break then. We'll take a ten-minute break and we'll come back. Just again to -
 24 hold on a second. We will take a ten-minute break. We will
- 25 come back and finish, at least the direct with Dr. Brookes, and

then we will see where we are.

- 2 MR. TYLER: Thank you, Your Honor.
- 3 **DEPUTY CLERK:** All rise. This Honorable Court now
- 4 stands in recess.
- 5 (Whereupon, a recess was taken from 10:48 until 11:00
- 6 a.m.)

- 7 DEPUTY CLERK: All rise. This Honorable Court
- 8 resumes in session. The Honorable Theodore D. Chuang
- 9 presiding.
- 10 **THE COURT:** Thank you, everyone. Please be seated.
- 11 Go ahead. Continue.
- 12 MR. TYLER: Thank you, Your Honor.
- 13 BY MR. TYLER:
- 14 Q. Now, Dr. Brookes, turning to some vignettes in the other
- 15 reports without going through each ad nauseam. Let's turn to
- 16 Exhibit-5A for a minute.
- 17 Dr. Brookes, can you tell us which report you see on your
- 18 | screen there?
- 19 **A.** This is report number 4.2 regarding series of images from
- 20 | figure two of a grant proposal.
- MR. TYLER: Ms. Cornich, if we can go to the next
- 22 page, please.
- 23 BY MR. TYLER:
- 24 | Q. Dr. Brookes, can you tell me what is the analysis that you
- 25 have done here?

1 A. So this is a reverse analysis tracing the final published

2 | image that you see in the top right corner. It's a series of

two Western blots side by side tracing that to a white box

4 | image. You will notice the final published image over on the

5 | right has 20H7. That is the name of an antibody that was used

in this Western blot.

In addition, you will see the title of the white box image contains the same text and, in fact, the handwriting in the top left corner of the piece of film also says 20H7. The file name of the white box image also contains a number 11 like many of the other images. The Western blots themselves are reporting levels of protein in various different patient samples. AC here is Alzheimer's Disease. YCI is young cognitively impaired, so these are patients of different stages of the Alzheimer's process.

In the image on the left, we have highlighted the red zone. The white box and enlarged that and moved it down here below the main image, and as you can see from the areas highlighted in blue, the pattern of bands is reproduced between the white box image and the final published image which leads to the conclusion that the white box image was the source. In addition to the bands themselves, you can see a number of noise features, most prominently this large vertical smudge below on the AD band is reproduced as well.

Q. And, Dr. Brookes, can you see any molecular weight markers

1 | in the white box image?

- 2 A. No. Despite the appearance of molecular weight markers on
- 3 | the left in the final published image, there don't appear to be
- 4 any molecular weight markers on the white box image.
- 5 Q. If we can go to the next page after you sort of made this
- 6 match. What are we looking at here?
- 7 **A.** This is essentially the ORI droplet applied to the
- 8 | left-hand set of blots and on the left slide is the same
- 9 analysis applied to the right-hand set of blots.
- 10 MR. TYLER: Ms. Cornich, if we can go to the next
- 11 page.
- 12 BY MR. TYLER:
- 13 Q. Dr. Brookes, what do we see here?
- 14 **A.** So this on the right is the white box image, as you just
- 15 | saw in the previous slides. On the left is a related raw
- 16 | image. You will note the raw image is missing the number 11
- 17 | from the file name. However, a number of other features are
- 18 common between the raw image and the white box image. The
- 19 | handwriting 20H7 in the top left corner, as well as all the
- 20 noise features that are shown in red here are common. The
- 21 | right-hand image has been rotated slightly. It's at a slightly
- 22 different angle, but, nevertheless, the noise pattern is
- 23 reproduced and so this leads to the conclusion that the raw
- 24 | image shown here is the digital source for the white box image.
- 25 **Q.** If you go to the next page. Dr. Brookes, can you tell us

what analysis you are doing here on this page?

- 2 A. So this is a forwards analysis. This is, as was
- 3 described, at the top is the raw image. At the bottom is the
- 4 | white box image, the relevant parts of both. In the middle is
- 5 where I have taken the raw image and subjected it to a
- 6 brightness adjustment in order to see if the bands in the white
- 7 box image appear. What you can see in red, the reason why this
- 8 particular 80 percent brightness was chosen is because this was
- 9 the adjustment level that lead to the appearance of the noise
- 10 that is in the red box.
- So you see in the two red boxes here that the noise in the
- 12 adjusted image and the white box image are identical, so
- 13 certainly the noise in the white box image appears to come from
- 14 the raw image. However, if you look at the areas highlighted
- 15 | in blue, you can see that the bands in the white box image do
- 16 | not appear in the raw image or in the adjusted version of the
- 17 | raw image.

- 18 **Q.** When you did that adjustment, were you at any point able
- 19 to make those bands appear?
- 20 **A.** What is shown here is an 80 percent brightness, but I
- 21 | attempted any and all combinations of brightness and contrast.
- 22 No adjustment was performed which was able to reproduce and
- 23 | cause these bands in the white box image to appear.
- 24 \mathbf{Q} . What conclusion were you able to draw by the fact that the
- 25 | red boxes match?

- 1 A. The red boxes matching essentially says these two images
- 2 | are digital relatives, that one is the source for the other
- 3 because the fingerprint of that noise is reproduced between
- 4 them, and that is also seen in other parts of the image such as
- 5 the handwriting in the top corner of both images.
- 6 | Q. Thank you. Let's move to maybe quickly through another
- 7 example of this. If we could bring up Exhibit-7A.
- 8 Dr. Brookes, what report is this?
- 9 **A.** This is report number four dealing with figure four from
- 10 one of the N.I.H. grant proposals.
- MR. TYLER: And if we can go to the next slide, Ms.
- 12 Cornich.
- 13 BY MR. TYLER:
- 14 Q. Dr. Brookes, did you, generally speaking, explain what the
- 15 | red arrows here are indicating?
- 16 **A.** Yeah. My apologize for the number of arrows on this
- 17 | slide. The final published image is in the center on the
- 18 | right-hand side of the slide. So this here is the final
- 19 published image. On the left is the corresponding white box
- 20 | image. You will note, again, number 11 in the file name. The
- 21 upper portion of this white box image in the top left, that is
- 22 | highlighted in red and then shown expanded and enlarged below
- 23 the final published image, so you can see from the matching
- 24 blue boxes here and here that this image is the source for
- 25 those particular bands in the final published image and then

1 | flipping things around, the bottom left, if we take the bottom

- 2 | left white box image that's highlighted in red, we take that
- 3 and blow it up and put it above the final finished image, you
- 4 can see that the other bands highlighted in blue here and here
- 5 | are also matching.
- 6 So this particular white box image which has a number of
- 7 | white boxes on it is the digital source for this final
- 8 | published image. And you can see a number of noise features as
- 9 | well. For example, this smudge here is reproduced in the final
- 10 image as well.
- 11 **Q.** And did you also do the droplets overlay analysis on this?
- 12 **A.** Yes. That was performed on this analysis as well and
- 13 | showed the same conclusions.
- 14 | Q. I think we can skip through and go to Page 5 of this or
- 15 | Page 6. I apologize.
- So, Dr. Brookes, after having done that analysis, what are
- 17 | we looking at here?
- 18 A. So this is, again, a forwards analysis. On the top is the
- 19 | raw image. On the bottom is the white box image trying to test
- 20 the hypothesis that the raw image can be adjusted in a way that
- 21 | makes the bands in the white box image appear. Again, as you
- 22 see in red here and here, there are a number of noise features
- 23 and background features which do appear to transpose from the
- 24 | raw image to the white box image. Notably, there are, in fact,
- 25 | a pair of bands which match up to a pair of bands on the raw

- 1 image and they, indeed, show up in the white box image.
- 2 However, as you can see, nothing in blue is in common between
- 3 | these images. No adjustment of any kind was able to make the
- 4 bands appearing in blue in the final image appear.
- 5 Q. Taking all that together, what were you able to conclude
- 6 about the relationship between these images?
- 7 **A.** So they are digitally related. The raw image appears to
- 8 be the parent of the white box image and the bands in the white
- 9 box image, however, don't have any provenance in the raw image.
- 10 **Q.** Is that true regardless of what different types of
- 11 | contrast and brightness adjustments you did?
- 12 **A.** Correct. Any and all adjustments failed to yield the
- 13 bands in blue.
- 14 Q. Let's now move to Exhibit-11A. Dr. Brookes, can you tell
- 15 us what figure this relates to?
- 16 **A.** This is report number eight regarding figure seven from
- 17 | the N.I.H. proposal.
- 18 **Q.** And then if we could fast forward to I guess page -- the
- 19 next page here.
- 20 Dr. Brookes, can you tell us what we see here?
- 21 | A. So this is a series of Western blots for one, two, three,
- 22 | four, five, six, seven different proteins across a number of
- 23 different conditions. The proteins are listed on the right.
- 24 PLC1 and NOS-PYK, PSD95. Over on the left are the purported
- 25 | molecular weights of those proteins. Down at the bottom is a

Western blot for a protein called NR1. In this case, that
protein is being used as a control. Fairly often when doing
Western blotting of a number of different proteins we Western
blot what is called a housekeeping protein or a protein that is
very common in cells as a way of ensuring the same amount of
protein has been loaded in each lane.

So it's fairly common when we want to quantify, for example, the amount of PSD95 or PKC, we would express the density of those bands relative to the density of the NR1 band as a way of normalizing that density to the amount of protein that is loaded in that particular lane.

- Q. Looking at the third and fourth rows there, could you just identify for us what molecular weight we are looking at for those two proteins?
- A. Yes. So on the top is a protein called PYK2. That has a native molecular weight of about 114 kilodaltons, so as you can see, it runs ever so slightly above the 100-molecular weight marker. In addition, PSD95, as the name suggests, has a molecular weight of 95 kilodaltons, and so that runs ever so slightly below the 100-molecular weight mark.

I found it surprising that it was possible to blot the two of these on the same gel because, in fact, they are barely separated. One is right above one of the markers and one is right below it.

So, if you were to run these on the same gel, they would,

1 in fact, appear very very close together, almost overlapping.

- 2 Q. If we could go to the next page. Dr. Brookes, could you
- 3 | just walk through what we're looking at here and what analysis
- 4 you did?
- 5 **A.** So these are the white box images which corresponds to the
- 6 published image that you just saw. Drawing your attention over
- 7 to the right, this is the Western blot in which PYK and PSD95
- 8 were run on the same gel. What is surprising about this is the
- 9 degree of vertical separation between the bands. You know,
- 10 there is not a lot of different between 114 and 95. And as I
- 11 | mentioned, you typically want to run those on separate blots.
- 12 So the fact that they both appear on this single separated by a
- 13 | very very large amount does not seem feasible or physically
- 14 possible for those bands to be real.
- 15 Q. In terms of the numbers that are counted out, what is the
- 16 relevance of those?
- 17 | A. Yes. So, as I mentioned, when you want to quantify the
- 18 data from this type of blot, it's typical to normalize it to
- 19 the amount of NR1. So, if you look at the two gels on the
- 20 right and count the lanes from left to right, you can see that
- 21 | they both have 13 lanes. However, if you look at the NR1 blot
- 22 over on the left, you can see that because it's handwritten NR1
- 23 | control. There are only 12 lanes. So that means for at least
- 24 one of those samples it would be impossible to calculate the
- 25 density relative to NR1 because you only have 12 NR1 bands to

normalize 13 bands of the protein of interest.

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Another interesting feature about these two blots on the right, they are ten well gels. There are ten lanes. And as you see here, all ten lanes have been used for loading of biological samples. That does not leave any room for loading a molecular weight marker and, in addition, there are no molecular weight annotations anywhere on these Western blot images which, again, makes it unusual that the final published image has molecular weight markings alongside the blots.

- 10 **Q.** Dr. Brookes, were you able to track down any data associated with this particular image?
- A. Yes. We were able to locate a PowerPoint file and the
 PowerPoint file contained a graph that was published alongside
 these Western blots and right clicking on the graph was able to
 bring up an Excel sheet and we were able to analyze the Excel
 sheet.
- Q. And with respect to the 12 versus 13, did you notice anything on that?
- A. What was unusual is that the NR1 data in that data set,
 there were 13 possible conditions, even though there were only
 label{eq: 12 bands} to see where were 13 sets of data in the spreadsheet so
 it's hard to see where at least one of those data sets came
 from given that there were only 12 bands to base the
 densitometry on.
- 25 Q. Thank you. Moving ahead to the next example. If we can

go to Exhibit-12A.

- 2 Dr. Brookes, can you tell us which analysis we are looking 3 at here?
- 4 A. This is report number nine looking at figure nine in the 5 N.I.H. grant.
- 6 MR. TYLER: If we could go to the next page, please
- 7 Ms. Cornich.
- 8 BY MR. TYLER:
- 9 Q. If you could us what we're looking at here?
- 10 A. So the top image is a white box image. It is a zoom in on
- an area of a white box image and in red, I've highlighted a
- 12 | series of six Western blot bands. Below in blue and orange is
- 13 | a false color gradient map applied to that in order to
- 14 highlight the edges of those bands and their shape and size.
- What's unusual about this, as you see underlined in red,
- 16 three of the bands appear very very similar shape, almost
- 17 | identical. The other three bands highlighted in white also
- 18 have a very unique shape that is reproduced across all three
- 19 bands. It's quite common in a Western block, for example, to
- 20 | see one band with a small indentation in it or a small smudge
- 21 on the right-hand side as you see here. It is highly unusual
- 22 to see several bands all of which appear to be identical, and
- 23 | it's just completely unheard of to find sets of bands, families
- 24 of bands that appear to be reproduced.
- 25 \mathbf{Q} . Have you ever seen that happen in one of your blots?

1 A. No. That is not something I have ever seen in a normal set of Western blots.

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- Q. Could you explain what the bottom, each of the bottom two figures are?
- 4 5 So in gray, this is a forensic image analysis tool known as error level analysis. Essentially when a JPEG image is 6 7 created, if the parts of that image all have the same 8 provenance, then the error in that image should be evenly 9 distributed. However, the ELA here, as you see in highlighted 10 by the red arrows, there appears to be a rectangular portion of 11 this image which has a different error level in the JPEG and 12 the rest of the image. And that is highly suggestive that this 13 part of the image, in fact, came from somewhere else. 14 we don't even need to do ELA image analysis to show this. 15 we look in green, this is just a gradient map applied to the whole image in order to extenuate the background noise, and you 16 17 can see the, generally speaking, the whole of the background of this gel image is green with yellow speckles. However, the 18 19 area highlighted, the rectangle in green, has significantly less yellow. It appears to be on a different background than 20
- Q. Do you remember ever seeing anything like this in one of your own blots?
- 24 **A.** I have never seen this in any normal Western blot.
- 25 Q. Now, let's turn to Exhibit-15A.

the rest of the image.

Dr. Brookes, can you tell us what we are looking at here?

- A. So this is a report looking at a number of different images and grant proposals and tracing them back to single source images.
- Q. Dr. Brookes, could you --

6 MR. TYLER: If you could go to the next page, Ms. 7 Cornich.

THE WITNESS: Okay. So in the top right, you see figure four of one of the ROR44 grant applications. The Western blot of interest in this case is highlighted in red. That is a Western blot for the nitrotarazine form of tau. That is what NY stands for.

Over on the left is a related white box image. You will note like all of the other white box images, it has 11 at the end of the title. In the top left is a white box area from that blot. We have enlarged that in red and brought it down underneath the enlargement of the NY tau blot from the figure. You can see not only the density of the bands are the same, they also have exactly the same relative position. Some are slightly higher than others. In addition, the width of the bands is the same. The left-hand band in particular appears slightly narrower than the other bands, and then there is a smudge to the upper left corner which is also reproduced.

So this, along with the ORI droplet analysis which I believe is on the next page of this report, shows that these

1 images are, in fact, the same image.

- 2 Q. And then if we go to the next page after that, what are we
- 3 looking at here?
- 4 A. So in C in the top right this is figure 13 from a
- 5 different R01 -- sorry -- R44 grant proposal. This shows a
- 6 Western blot for the protein Beta Actin and below the bands you
- 7 can see a prominent pattern of smudging immediately below the
- 8 bands on the right.
- 9 On the left is a white box image. It appears to have a
- 10 similar file name to the other white box image. However, now
- 11 instead of 11 at the end, it has 1232 at the end. This white
- 12 box image has an area of the upper left blot that has been
- 13 taken in red and enlarged and moved below the published image.
- 14 And as you can see, the images are the same. The bands
- 15 are the same. The smudges above and below the bands are the
- 16 same. And that's also reproduced in the ORI droplet analysis
- 17 as well.
- 18 | Q. If we can go to the next slide, is this that droplet
- 19 | analysis?
- 20 A. Correct.
- 21 | Q. Now, let's go to the last slide. Dr. Brookes, if you
- 22 | could explain what we can see now here between all three of
- 23 | these figures?
- 24 | A. Okay. So on the right you see the two white box images
- 25 | for the two completely separate proteins in completely

different figures. And you see on the left is the raw image.

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What's unusual here is if we actually count up the bands and we look at the figure legends from the figures reporting on what are the treatments in each group, and so focusing on the middle, if we look at the middle blot, if we count six bands over from the left, we can see that the six band corresponds to treatment of the sample with a thousand units of the drug.

So lane number six contains the sample that was treated with a thousand units of drug. If you now go to the white box image on the right and count over six bands to here, you can see that that exact same lane in the exact same position of the gel now corresponds to a hundred units of the drug.

You can't load more than one biological sample in the same lane on a gel. So this reporting of what has apparently happened in the samples of these drugs is impossible.

- Taking also a step back, Dr. Brookes, is it possible for the raw image to be the parent for both the figure four in the middle and the figure 13 on the right?
- 19 The raw image has the same handwriting, the same 20 fault, the same blotches. In addition, one particular 21 alternative explanation we thought about here was the 22 possibility for stripping and reprobing because we are looking 23 at two different Western blots for different proteins. 24 However, as mentioned, when you take a Western blot membrane 25

out and wash it and strip it and put another membrane and

- 1 another antibody and put it in a new file wallet, there are
- 2 different bubbles and different bits of hair and whatever gets
- 3 | in there. It is impossible that the exact same pattern of
- 4 | blotches and noise and bubbles and everything would appear on
- 5 all three. The only possibility here is that these images are
- 6 | all digital relatives of each other.
- 7 Q. And with respect to the raw image, can that have been the
- 8 source of the bands for figure four and then a different set of
- 9 bands for figure 13?
- 10 **A.** The bands in the white box image do not find provenance in
- 11 the bands in the raw image. There are unusually -- unlike many
- 12 of the other raw images in this case, there are some bands in
- 13 this raw image. However, they do not match the bands in the
- 14 white box images or the published images in terms of the slope,
- 15 the shape, the size, the spacing, the distribution and the
- 16 density.
- 17 | Q. Let's now turn to Exhibit-8A. Dr. Brookes, can you tell
- 18 us what report we're looking at here?
- 19 **A.** This is report five. This is looking at figure 14 from
- 20 the R44 proposal.
- 21 | MR. TYLER: Ms. Cornich, can we go to the next page?
- 22 BY MR. TYLER:
- 23 \mathbf{Q} . What do we see here with this analysis?
- 24 \mid **A.** In the top right is the final published image. We are
- 25 looking at three different proteins, TLA -- TLR4, Filamin A and

1 the alpha 7 nicotinic acetylcholine receptor. Over on the left

2 is a white box image. You will note from the file name that

the three proteins that we just mentioned are listed in the

4 file name along with 112 appended to the end of the file name.

So, if we look at the white box image, we have highlighted that in red and bring it down and put it below, the proteins are in a different order than they are vertically, so in the final published image Filamin A is at the bottom, whereas in

9 the white box image it's at the top. That does not really

10 matter. What is important is the bands match. Everything is

11 | the -- the areas in blue, the bands in the white box image are

12 the bands in the final.

MR. TYLER: Ms. Cornich, if we can go to the next

14 slide.

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15 BY MR. TYLER:

- 16 **Q.** Dr. Brookes, now what do we see here?
- 17 **A.** So this shows the corresponding raw image. At the top
- 18 that is the raw image. The white box image is in the middle.
- 19 You can tell that they're related by the handwriting on both
- 20 | images as well as the file names minus the 112 part. And then
- 21 | at the bottom we have a very unusual image which is part of the
- 22 same digital image family. It has the same handwriting. But
- 23 | in this case there are very solid black bands and there does
- 24 | not appear to be a white box.
- 25 \mathbf{Q} . Just to be clear, this is what you are talking about when

you are talking about --

- 2 A. Correct. Yes. This -- this image appears to contain a
- 3 | number of features and bands which are not present in the raw
- 4 image.

- 5 MR. TYLER: Ms. Cornich, if we can go to the next
- 6 page.
- 7 BY MR. TYLER:
- 8 Q. What are we looking at here, Dr. Brookes?
- 9 A. So this is comparing the pattern of bands in the white box
- 10 | image with the pattern of bands in the extra image that we saw
- 11 at the bottom of the page previously. Again, a number of noise
- 12 | features are carried over. The arrow over on the far right
- 13 indicates a small vertical blemish to the top left of the band.
- 14 | That appears in both images. If we count a couple of bands
- 15 over, on the left here, you can see there is a band that
- 16 appears to have its bottom right corner sheared off as
- 17 | indicated by the red line. And then the arrow in the middle of
- 18 | the image there appears to be an indentation in the top right
- 19 | corner of this band.
- 20 So these bands essentially overlap and this leads to the
- 21 | conclusion that these solid bands in the third image were the
- 22 source for the bands in the white box.
- 23 **Q.** If we can go to the next page?
- 24 | A. So this shows an ORI droplet of those same bands. What's
- 25 important to realize here is not just each individual band.

1 It's the size, the shape, the slope, the positioning, the

- 2 spacing between the bands. All of that is also the same.
- 3 MR. TYLER: Ms. Cornich, if you can go back to the
- 4 previous slide and blow up the bottom of this one.
- 5 BY MR. TYLER:
- 6 Q. Dr. Brookes, is there anything different that you observed
- 7 about the resolution here?
- 8 A. So these bands are, indeed, very unusual looking.
- 9 Typically when we see bands, the amount of noise in an image
- 10 | should be uniform across the image. What you are looking at
- 11 here is, indeed, there are blotches of noise and pixilation
- 12 and, you know, essentially fuzz in the background of the image.
- 13 You know, the little spots here and there. The bands, however,
- 14 | are very unusual because they appear to be sharp. They are
- 15 very sharp edges. And that is highly unusual to find a
- 16 different resolution. They almost look as if they are
- 17 different resolution. It's highly unusual to find bands of a
- 18 different resolution come against a generally noisy, fuzzy,
- 19 | blotchy background.
- 20 **Q.** When you say "highly unusual," is that something you ever
- 21 | observed in your own blots?
- 22 **A.** That is not something I ever observed in my own blots or
- 23 | anybody else that I know.
- 24 MR. TYLER: Ms. Cornich, if we can go to I believe
- 25 | it's Page 6 now.

BY MR. TYLER:

- 2 Q. Dr. Brookes, can you tell us what we see here on this
- 3 page?

- 4 **A.** So this is a type of analysis which was described in the
- 5 original pipeline. This is a histogram analysis, so what this
- 6 is doing is basically measuring the number of black and white
- 7 and all the shades of gray in between, the number of pixels at
- 8 each shade.
- 9 So, looking over on the left is this extra image. If we
- 10 look here, we have taken a section of that image in red and
- 11 blown it up and put it in the middle here. And what I have
- 12 done now is to highlight in blue some normal looking Western
- 13 blot bands. These are -- this is what Western blot bands look
- 14 like. They are black in the middle. They fade at the edges.
- 15 They're fuzzy as you get further out from the center.
- So, in Photoshop what can be done is a histogram analysis
- 17 | and so over on the right this is a graph showing on a scale
- 18 | along the bottom, black on the left and gray and white on the
- 19 | right, gray in the middle showing the distribution of different
- 20 pixel intensities.
- 21 So that's a very common standard looking pixel intensity
- 22 histogram. In contrast, if we then go to the unusual looking
- 23 | bands, these are highlighted in blue here and the histogram of
- 24 those is shown in the top right. And what I would like you to
- 25 | focus on is this right here. There is an unusual concentration

- of pure black pixels in this histogram. This is not seen.
- 2 This is highly unusual. This would be seen if, for example, I
- 3 were to add bands digitally using pure black color. This is
- 4 | not seen for natural looking bands as you see by comparison
- 5 with the histogram below.
- 6 Q. When you say it is not seen, have you ever seen that in
- 7 any of your blots?
- 8 A. I have seen that in other analyses that I have done of
- 9 other suspect images outside of this case. I have used
- 10 | histogram analysis to actually show that certain bands are not
- 11 part of a block and have a different profile and came from
- 12 somewhere else.
- 13 Q. But in your own blots --
- 14 | A. But in my own blots, I haven't seen this.
- MR. TYLER: Ms. Cornich, if you could bring up
- 16 Exhibit-23.
- 17 BY MR. TYLER:
- 18 \mathbf{Q} . Could you just tell us what this is, Dr. Brookes?
- 19 **A.** This is a description of an analysis method that was
- 20 applied called a terminal digit analysis in report number 5.1.
- 21 | Do you need me to go in to what was found or --
- 22 Q. Yeah. If we can go to the bottom and get the last
- 23 | paragraph of Page 2. First of all, can you explain what
- 24 | terminal digital analysis is?
- 25 **A.** Okay. So, when numbers are generated from randomly

Case 8:24-cr-00211-TDC Page 91 of 206 occurring stochastic processes, such as biological experiments, 1 the last digit on the right has an equal probability of 2 occurring across the range. So between zero and nine, one will 3 be the last digit, the exact same number of times that two or 4 5 three or four or five or six will be the last digit. So, in a terminal digit analysis, we basically take all of 6 7 the numbers in a data set, and so in this particular example you can see there are 3,208 numbers. We would then expect each 8 9 terminal digit to appear 320 times. The actual number of 10 appearance of the terminal digit is shown in the third column 11 right here. 12 As you can see, for example, number seven appears more 13 often than expected and number nine appears way more often than 14 expected. We can perform a statistical test to compare the 15 observed versus the predicted occurrence. And the value that is reported in red is essentially a probability. 16 17 probability that this set of numbers arose from a random stochastic process such as a biological experiment. 18 19 That probability in this case for this data set is one in 20 a hundred million. There is a one in a hundred million

likelihood that these numbers came out this way from a random stochastic process.

21

22

23

24

- Just to circle back, what were the data points? Where did they come from?
- Α. These were densitometry data for Western blots in the

Journal of Neuroscience paper.

- 2 Q. Dr. Brookes, in the course of your work on this case, did
- 3 | you ever find a raw image that matched up to any bands --
- 4 matched up to the bands of the white box or published image in
- 5 | the images that you looked at?
- 6 **A.** No.

- 7 Q. What did you conclude -- we didn't cover every report here
- 8 during your testimony. But what did you conclude about the
- 9 various figures we reviewed in terms of the figures matching
- 10 | the research record?
- 11 A. I concluded overall that there were numerous examples of
- 12 | bands appearing in the white box and the final published
- 13 | versions of images that have absolutely no provenance
- 14 | whatsoever in the underlying raw data images.
- I also concluded that there were problems with the
- 16 statistics from the densitometry and the data presented in the
- 17 | graphs. There were also a number of just physical
- 18 | impossibilities, for example, regarding the pH scales for the
- 19 | isoelectric focusing, regarding having more than a hundred
- 20 percent of a protein being in a particular conformation on the
- 21 | Excel spreadsheet and then having, for example, number nine
- 22 appear way more often than is statistically probable in the
- 23 | final data set.
- 24 So my general conclusion is that on numerous occasions
- 25 here the data has been fabricated.

1 Q. Does that include reports that we covered today and also

- 2 ones that you entered and the exhibits that we did not cover
- 3 today?
- 4 A. Yes. Everything is entered into evidence.
- 5 Q. With respect to all the images that you conclude were
- 6 fabricated, did any of them lead to the weakening of the
- 7 | conclusion in the grade application or article with respect to
- 8 PTI125 or its diagnostic companion?
- 9 A. No. Frequently the alterations that were done, that
- 10 appear to have been done to the images strengthen the
- 11 | conclusions.
- MR. TYLER: Your Honor, if I may have one moment?
- 13 **THE COURT:** Okay.
- 14 MR. TYLER: Nothing further, Your Honor.
- 15 THE COURT: Okay. Before we go to cross-examination,
- 16 | I normally wait until I get until the end of all testimony to
- 17 | see if I have any questions, but there is one I just want to
- 18 make sure I understand that will help me with even the cross.
- 19 I don't think you really explained what a white box image
- 20 is and where it comes from. Maybe you can just clarify that
- 21 | for me before we get to the cross-examination.
- 22 **THE WITNESS:** Okay. So I was provided with sets of
- 23 | images by Agent Weeks. I was provided typically with a raw
- 24 | image and a corresponding white box image. I say corresponding
- 25 because they essentially have the same file name and share a

number of features. I don't know where those images came from. 1 What is a white box image? Is that an --2 THE COURT: 3 THE WITNESS: White box is just the terminology that I applied to those images to distinguish them from each other. 4 5 The origin of the white box is unknown to me. I'm just trying to even understand what 6 7 we are talking about. Are you saying there was an image that 8 instead of having this sort of grayish background it was like a 9 letter box just focused on the bands? 10 THE WITNESS: One area. Yes. Sorry, Your Honor. 11 I'm interrupting you. 12 THE COURT: Is that a normal thing to do in science, 13 generally, is you might generate a white box to focus on 14 certain areas or not? 15 THE WITNESS: That breaks the fundamental rule of 16 image analysis which is that you have to treat the whole image 17 equally, so it's highly unusual to find a white box that --18 which appears to be superimposed upon the raw image in that 19 way. 20 THE COURT: Okay. I think I understand, but we will 21 go to cross-examination now. Thank you. 22 Just to clarify, Mr. Tyler, Exhibits-50 and 52, when those 23 came up, so those are in evidence for purposes of the hearing.

Any issues with that from either side? Fifty and 52?

MS. BEIDEL: No, Your Honor.

24

```
MR. TYLER:
                            No, Your Honor.
 1
 2
               THE COURT:
                            Go ahead, Ms. Beidel.
 3
               MS. BEIDEL:
                             Perhaps starting with exhibits first.
    If it's acceptable to the Court, I could let you know which
 4
 5
    exhibits I would intend to offer and see if the government has
    any objection.
 6
 7
               THE COURT:
                            Sure.
                             So Exhibit-87 to 87-6, 91 and 91-1 and
 8
               MS. BEIDEL:
 9
    then everything from 101A through 137 and 146 through 161.
10
               MR. TYLER:
                            I'm sorry. We are trying to -- can you
11
    repeat that again?
12
               THE COURT:
                            Just give us the range one more time.
13
    Sorry.
14
               MS. BEIDEL:
                             87 to 87-6, 91 and 91-1, 101A through
15
    137 and 146 through 161.
               THE COURT:
16
                            Okay.
17
               MS. BEIDEL:
                            May I proceed, Your Honor?
               THE COURT:
                            Any issues with those, Mr. Tyler?
18
19
                            If you can just give me one moment, Your
               MR. TYLER:
20
    Honor.
21
               THE COURT:
                            Okay.
22
               MR. TYLER:
                            Your Honor, we have no objection with the
23
    exception of the video clips because we just haven't had a
    chance to review those yet, but everything else we are fine.
24
```

What numbers are those?

25

THE COURT:

135 through 137. 1 MR. TYLER:

2 THE COURT: Okay. Are you going to use those in any

3 great way?

4

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MS. BEIDEL: No. That's fine, Your Honor.

> THE COURT: Okay.

MS. BEIDEL: I can skip that.

> THE COURT: Okay.

8 CROSS-EXAMINATION

9 BY MS. BEIDEL:

- 10 Good morning, Dr. Brookes.
- 11 Good morning.
- 12 I want to start where we stopped which was you were
- 13 discussing -- you first said alterations that were done and
- 14 then you corrected yourself to alterations that appear to have
- 15 been done. Tell me why you made that distinction.
- 16 I'm basing my conclusions on the appearance of what was
- 17 presented to me.
- And it's correct that you don't know for certain that 18
- those alterations were made in the way that you testified to; 19
- 20 correct?
- I don't know how the alterations were made to the images. 21
- I know that they were made digitally for the reasons described, 22
- but how digitally, I don't know. 23
- 24 It's your testimony under oath today, as you sit here,
- 25 that you are certain that there were digital alterations to

1 these images?

- 2 A. Correct. Yes.
- 3 **Q.** You could not be wrong in the way that you were wrong when
- 4 you accused your colleague at the University of Rochester of
- 5 manipulation?
- 6 A. I do not believe I am wrong when I testify that these
- 7 | images were digitally altered.
- 8 Q. And isn't it true that how you learned you were wrong in
- 9 that case was that that colleague was given the opportunity to
- 10 provide you with an original that convinced you that you were
- 11 wrong?
- 12 **A.** The colleague came to my office, yes, sir.
- 13 **Q.** And in this case, you haven't had any conversations or
- 14 other interactions with Dr. Wang through which he would have
- 15 the opportunity to provide you with such originals; correct?
- 16 A. I have never met Dr. Wang before.
- 17 Q. So, all you have is the record provided to you by Special
- 18 Agent Weeks?
- 19 **A.** That -- yes, that is correct.
- 20 **Q.** And if there were, say, other originals that showed the
- 21 | provenance of the bands that you analyzed, then your opinions
- 22 | could change; correct?
- 23 | A. If those original images exist, it still would not change
- 24 | the fact that the bands do not exist in the raw image, so it
- 25 | would be unlikely to change my opinion. It would depend on the

- 1 data that was presented, if such images do, indeed, exist.
- 2 Q. You connected the raw images to the white box images
- 3 essentially by assuming that the only way that those white box
- 4 images could exist there is if they were a component of that
- 5 original image; is that fair?
- 6 A. No. The white box, as I believe was just discussed with
- 7 His Honor, my thought is that the white box is a digital
- 8 addition to the raw image. The white box was not an original
- 9 component of the raw image as you just stated.
- 10 Q. So, in other words --
- 11 **A.** Because if it was, then it should be possible to
- 12 | manipulate the raw image to make the white box appear and, as
- 13 was described, that is not possible.
- 14 **Q.** So you can see that the white box images are cut and
- 15 pasted on top of the supposed raw images; correct?
- 16 | A. I don't know how the white box or the bands within the
- 17 | white box came to appear on the raw image. All I know is that
- 18 | the information; namely, the bands, is not present in the raw
- 19 image.
- 20 **Q.** Well, didn't you just testify that it's impossible to make
- 21 | the white box image by altering the raw image?
- 22 **A.** It's impossible to make it by altering the raw image using
- 23 | acceptable image manipulation standards such as altering the
- 24 brightness of the -- or the contrast of the whole image.
- 25 \mathbf{Q} . So one possibility is that the white box image is cut and

- 1 pasted on top of the supposed raw image; correct?
- 2 A. That is a possibility. I don't know why anybody would do
- 3 that. It's not a standard scientific practice.
- 4 Q. So, if the white box image were cut and pasted on top of
- 5 the raw image, then all of your analysis talking about the
- 6 other things and the raw image outside of the white box are
- 7 | irrelevant; correct?
- 8 **A.** No.
- 9 Q. Well, if the white box image is the only image that lead
- 10 to the figure, then if you are talking about sourcing the rest
- 11 of the image to the raw, that does not matter?
- 12 **A.** The bands in the white box image are of the data. And if
- 13 the white box image was pasted in from another image, albeit
- 14 unknown at this stage, then that other image is the raw data,
- 15 | in which case given that nobody has seen those other images
- 16 that by your definition states that the raw data do not exist.
- 17 \mathbf{Q} . So would you agree with me that there exist some other
- 18 | image that contains the raw data or at least existed at some
- 19 | time?
- 20 **A.** There could be additional images that contain the pattern
- 21 of bands seen in the white box image. At no point during these
- 22 | investigations have I found such images.
- 23 | Q. But you can't testify with certainty as you sit here
- 24 today, that such images do not exist; correct?
- 25 A. I cannot say that they do not exist. What I can say is

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- that from a standard scientific practice, it would be expected 1
- that those images should be retained and when raw data is 2
- 3 requested, those should be the images that would be provided
- not, as you are attempting to say, some unrelated raw image 4
- 5 with splotches and no bands. If somebody asked me for the raw
- data, I would expect that I would provide them with the raw 6
- 7 data, not some unrelated image.
- 8 But weren't there two kind of ultimate rules about image
- 9 manipulation and presentation that come from this JCB paper.
- 10 Do I have that right?
- 11 Correct. Yes.
- 12 And is one of them what you just said?
- 13 There are two general rules. One is we shouldn't adjust
- 14 an image to change the informational content of that image.
- 15 And the other is that if provide -- if manipulating an image,
- we should adjust the whole image evenly to the same level. 16
- 17 So, for example, taking a raw image and pasting in another
- image that contains the white box bands, that changes the 18
- 19 informational content of the raw image so that breaks that
- 20 rule.
- Is there anything in the JCB paper about the number of 21
- 22 years one needs to retain that underlying data?
- 23 At the time those guidelines were written, which is I
- 24 believe 2006, I do not recall if there was anything in there
- 25 about data retention standards. As I'm sure you are aware, the

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- federal government has recently revised its standard regarding 1
- data retention policies relative to the funding period of a 2
- 3 grant. The typical expectation is six years after the grant
- has finished all the publications have been published. 4
- 5 So, for example, just using that six-year period, the
- paper that you referred to in Neurobiology of Aging was 6
- 7 published in 2017 suggesting the work was done previously;
- 8 correct?
- 9 Α. Yes.
- 10 And so we would be outside of the retention period of that
- 11 law image and data according to that statement; correct?
- 12 At the time I was presented with those figures for the
- 13 purposes of this investigation it was 2002, which would had
- 14 been within the statute of limitations for data retention.
- 15 I think you meant 2022. You said 2002.
- 16 Sorry. 2022. Sorry. That would be five years after that
- 17 paper was published. And, generally, in that role from the
- N.I.H. it's the latter of either when the paper was published 18
- or when the grant funding ended. 19
- So I don't know when the funding that was used to do those 20
- 21 studies ended relative to when the paper was published.
- 22 MS. BEIDEL: Your Honor, could I approach to get
- Exhibit-52? 23
- 24 THE COURT: Yes.
- 25 BY MS. BEIDEL:

- In your word, you said all of the bits. I want to talk 1
- about all of the bits of the experiment. 2
- 3 Okay. Α.
- That lead to ultimately the digital images that you 4
- 5 analyzed, so working backwards, there is digital images.
- just talked about how there could be potentially other digital 6
- 7 images that you didn't see; correct?
- 8 You correct.
- Then there is film that looks like this; correct? 9
- 10 That is the developed film. Yes.
- 11 Developed film. Did you review any developed film from
- 12 Wang's lab in connection with this case?
- 13 No. No developed films were provided.
- 14 And then using this apparatus there is a gel that's
- 15 transferred to a nitrocellulose membrane; correct?
- Uh-hum. 16
- 17 And did you review any nitrocellulose membranes or gels in
- connection with Dr. Wang's? 18
- 19 Typically gels are very fragile. They don't store well,
- 20 so it's fairly common practice after the transfer of the gel to
- 21 the membrane to throw the gel away. Sometimes we will stain
- 22 the gel to see if there is any residual protein left in the gel
- 23 to see if the transfer happened properly. The nitrocellulose
- membranes, typically those are kept for a week or two after. 24
- 25 The main reason we don't keep a lot of nitrocellulose around,

- it is explosive. And keeping large quantities of 1
- nitrocellulose in binders in the lab is not a very safe 2
- 3 practice. So the nitrocellulose members are also typically
- discarded a week or two after the blot has been developed onto 4
- 5 the film.
- So fair to say you didn't review any gels or 6
- 7 nitrocellulose membranes for this case?
- 8 That would be unusual.
- 9 Okay. You've talked a lot about Special Agent Weeks
- 10 providing you with selections of images I quess going back to
- 11 2022; is that right?
- 12 Correct. Yes.
- 13 Did you provide Special Agent Weeks with certain
- 14 parameters as to what images you wanted him to provide?
- 15 No.
- 16 He chose the images and sent them to you; correct?
- 17 Agent Weeks sent them to me in sets typically of three.
- 18 Here is a grant. Here is some images I would like you to
- 19 analyze relative to that grant.
- 20 Are you aware of whether Special Agent Weeks had any
- 21 particular methodology for choosing those images?
- 22 I don't know how those images were chosen.
- 23 You mentioned you didn't speak with Dr. Wang in connection
- 24 with this case. Did you speak with anyone else from his lab at
- 25 CUNY or anyone from Cassava, the company that he was working

1 | with?

- 2 A. No. Not at Cassava. I believe when the original
- 3 discoveries were made regarding the figures in the papers -- so
- 4 I got into this case because I was asked by a colleague to look
- 5 at a set of raw data that were provided by Dr. Wang to the
- 6 editors of the paper, of the Journal of Alzheimer's Disease and
- 7 Therapy. In doing that, I found misconduct evidence in the
- 8 preparation of those figures. I reported that. One of the
- 9 parties that I reported that to was the research integrity
- 10 officer at CUNY.
- 11 **Q.** Do you know whether CUNY concluded that there was any
- 12 research misconduct in connection with that allegation?
- 13 A. I don't know regarding that allegation. I know CUNY ran
- 14 | an investigation, and I believe it's still ongoing, so the
- 15 results of that are unknown to me.
- 16 Q. So, as we sit here today, CUNY has not made any final
- 17 | findings with respect to research misconduct on behalf of
- 18 Dr. Wang, as far as you know?
- 19 | A. To the best of my knowledge, CUNY's investigation is still
- 20 ongoing.
- 21 **Q.** Have you ever visited Dr. Wang's lab at CUNY?
- 22 **A.** Never.
- 23 **Q.** Have you in any way observed his Western blotting process
- 24 recorded or otherwise?
- 25 **A.** No.

- could have access to any hard copy files that might exist in 2
- Dr. Wang's lab? Films? Lab notebooks or the like? 3
- I believe I may have asked are there films, are 4
- 5 there lab notebooks.
- And were you provided with any? 6
- 7 If those materials exist, I was not provided with them.
- 8 I'm not aware of their existence. My analysis was solely based
- 9 on the digital files that were provided to me by Agent Weeks.
- 10 And those files were provided by e-mail, and then I
- 11 believe you said on a disk at some later point?
- 12 There were two DVD's, one of which contained JPEGs
- and another which contained PowerPoint files. 13
- 14 And I think your number was something like 2,300?
- 15 2,300 JPEGs. The original number was slightly higher.
- understanding is that disk contained JPEGs from a computer, and 16
- there were a number of images on the case including, for 17
- example, photographs and desktop backgrounds which were 18
- 19 irrelevant to the case, so when they were removed 2,300 is the
- number of JPEGs relevant to the case that were on that disk. 20
- 21 We've received something over a million records in
- 22 discovery in this case. You certainly did not review all of
- 23 that at any point?
- 24 In the case of the PowerPoints and the JPEGs, it was a
- 25 frequent practice of mine to have a window open on one side of

- my desktop with the enlarged preview feature open for the JPEGs 1
- and then a JPEG of interest in the case on the right and scroll 2
- through 2,300 images to see if I could find one that matched. 3
- Right. 4
- 5 And that was also done with the PowerPoints. In one case
- I sat for an afternoon and opened 2,900 PowerPoints in order to 6
- 7 look at the images within those PowerPoints to see if any of
- 8 them matched, so there was somewhat of a group-force approach
- 9 to finding matches.
- There is some delta, obviously, between that 2,300 plus 10
- 11 2,900 and the government produced records in this case and you
- 12 didn't have access to any of that; correct?
- 13 Α. No.
- 14 Have you ever reviewed any correspondence from Dr. Wang,
- 15 e-mails, phone records, voicemails, anything like that?
- None of those have been provided to me. 16
- 17 Did you do meta data analysis of the images that you
- analyzed? 18
- 19 Other than the title, no. One of the issues here is that
- 20 the files were sent by e-mail and so, for example, when I
- 21 received that file and then save it to my computer, now the
- 22 creation date of that file is when I saved it. So in many
- 23 cases the meta data would not had been any use to my analysis.
- 24 My understanding is that Agent Weeks has that data.
- 25 **Q**. But if he does have that data, you haven't analyzed it?

- I was not told anything about the meta data to provide
- 2 context to the images that I analyzed.
- 3 So, as you sit here today, you can't use a forensic meta
- data analysis traced backwards from the publication image in 4
- 5 the grant application, let's say, to the raw image that you
- analyzed? 6

- 7 The titles do provide some hints and in a couple of cases,
- regarding the PowerPoint files specifically, I was able to find 8
- 9 the creation date of the PowerPoint file and that matched
- 10 relative to the time frame in which a paper or a grant was
- 11 submitted. And that is in the reports textually.
- 12 Are you familiar with an MD5 hash value of a document?
- 13 That's not an analysis that was performed in this
- 14 case.
- 15 To the extent any of the images in this case have a
- unique digital fingerprint number, you did not perform an 16
- analysis of that to trace it back to the original file? 17
- No. Correct. 18
- 19 Okay. Now, you mentioned the naming convention where a
- lot of these files end in 11; correct? 20
- 21 Correct, yes. Many of the white box images ended in dash
- 22 11 or dash 112 or dash 1212.
- 23 And you also mentioned there could be other stripped and
- 24 reprobed versions of an image or perhaps different exposure
- 25 times of an image for the same experiment; is that right?

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We specifically ruled out stripping and reprobing as a 1 potential explanation for additional images because when that 2 3 happens, the blog comes out of the cassette, gets washed and gets exposed to another antibody and reexposed. The chances of 4 5 being able to line up the film in exactly the same position the second time around is zero. Everything is going to be off by 6 7 fractions of millimeters. The noise pattern will be different. 8 So we specifically ruled out stripping and reprobing as a 9 potential source of additional images because then the noise 10 pattern would not be the same between the raw and the white 11 box. 12 Setting aside that source question for a second, there are 13 certainly some laboratory practices when you are producing 14 Western blots that you could use the same set of samples and 15 the same gel or membrane and either do different exposure

letters, strip and reprobe it and probably some other options to create multiple film images?

So the example you have, the piece of film you have, that is actually two Western blots exposed four times, so we put it down in the cassette once, flip it around, do it again, turn it upside down, do it again and then flip it one last time. you will see that the bands in the four quadrants of that piece of film are a slightly different darkness or brightness because those are four different exposures, and I believe in the top corners of there you can see it says like ten seconds, 30

- The time of each exposure is written in the corners 1 seconds.
- 2 of the piece of film.
- So using the film in Exhibit-52, for example, there is 3
- four different exposure times of two --4
- 5 Of two Western blots. Α.
- So we have eight images? 6
- 7 Α. Correct.
- 8 And there is really no limit on how many different
- 9 exposures one could run to get to the ideal image; correct?
- 10 Correct. You can expose for ten seconds. You can expose
- 11 for five minutes. Generally, with exposures what tends to
- 12 happen is if you have very very low abundance proteins the
- 13 longer you go with the exposure you may, indeed, be able to
- 14 pick up the protein of interest, but the background noise is
- 15 going to become oppressive to the point where you can't
- actually see the bands because the whole membrane will turn 16
- 17 black.
- 18 So back to Dr. Wang's naming convention. You have some
- 19 without a terminal digit and some 11s. It's certainly possible
- 20 that there are images one through ten out there somewhere;
- 21 correct?
- 22 It's possible there are additional digital images, but
- 23 that still doesn't explain where the bands in the white box
- 24 image came from.
- 25 Did you do any investigation as to how Dr. Wang was taught

to do Western blotting? 1

- That's not material to my role here, how he was taught. 2
- Let's say, for example, he was taught not to use molecular 3
- weight markers, that is not something you would be aware of; 4
- 5 correct?
- That would be highly unusual and that would be bad 6
- 7 scientific training. If I may add for the record, the cost of
- 8 molecular markers, one tube with enough for a hundred
- 9 experiments costs about \$35, so this is not an expensive
- 10 burdensome addition to any experiment. This is something that
- 11 costs pennies which anybody running Western blots does by
- 12 default.
- 13 You are familiar with a publication or a set of
- 14 publications called Plus One; correct?
- 15 Plus One is a journal I believe. When was it founded?
- 20 -- late 90s. 16
- 17 MS. BEIDEL: Ms. Blackwood, could we show
- Exhibit-159, please? 18
- 19 BY MS. BEIDEL:
- 20 So you will see, Dr. Brookes, that this is an article from
- 21 Plus Biology called blind spots on Western blots, assessment of
- 22 common problems in Western blot figures and methods reporting
- 23 with recommendations to improve them. And the received date is
- 24 June 7, 2022.
- 25 Do you see that?

- Correct. Yes. Α. 1
- If you look in the abstract about midway down it says, our 2
- 3 data show that most published Western blots are cropped and
- blot source data are not made available to readers in the 4
- 5 supplement. Publishing blots with visible molecular weight
- markers is rare and many blots additionally lack molecular 6
- 7 weight labels.
- 8 Do you see that?
- 9 I see what that says.
- 10 Have you done an analysis of Western blots similar to this
- 11 paper to determine the prevalence of use of molecular weight
- 12 markers?
- 13 That has not been an area that I have actively researched.
- 14 But I would say that this statement here from the authors is a
- 15 critique. This is a statement of a problem in the scientific
- literature. So, regardless of whether something is common, 16
- 17 this is being flagged in this paper because it's being
- described as a lack or a problem. That problem may, indeed, be 18
- 19 very very common, but that does not make it not a problem.
- 20 I understand that. But wasn't it your testimony that
- 21 molecular weight markers are very frequently used, especially
- 22 in all the Western blots you've observed?
- 23 It was my testimony and I stand by that testimony.
- 24 Western blots, standard use of Western blots includes molecular
- 25 weight markers. This is talking about published Western blots,

1 so it can be very common, in fact, to use a Western blot

2 | molecular weight marker on the initial gel and then for

3 | whatever reason, maybe space constructions in the journal, to

leave off the molecular weight markers in the final published

5 version.

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Q. As you sit here today, can you direct us to a paper that requires the use of molecular weight markers in Western

8 blotting?

A. A journal that I'm on the editorial board of. The Journal of Molecular and Cellular Cardiology, JMCC. They have a series of data standards. That data standards essentially state that not only must you include molecular weight markers on your images, if you produce any Western blot in that journal, you are required to provide a supplemental data file that includes all of the original gel images with their molecular weight markers to the journal during the submission process.

And more and more journals are now adopting data transparency standards where it is common practice to not only demand the Western blots, but the original images with the full height gels and the molecular weight markers that led to those images.

- **Q.** That's for that particular journal?
- A. That's for that journal and I know of several others who have documented similar standards. This is an area where many journals now are becoming more stringent in the quality of

- Western blot data that they will accept for publication. 1
- Has the N.I.H. adopted that standard? 2
- 3 The N.I.H. does not publish papers, so they would not have
- need to. 4
- 5 So at the time that Dr. Wang performed his research, the
- N.I.H. did not have a particular standard requiring use of 6
- 7 molecular weight markers; isn't that fair?
- 8 That's fair to say. I don't know why the N.I.H. would
- 9 dictate that standard. Typically, N.I.H. would not dictate
- 10 standards about how individual experiments are supposed to be
- 11 I'm not aware that they do that for any other method
- 12 aside from Western blotting. They don't tell you how to use a
- 13 microscope, for example. They don't tell you how to do other
- 14 biological science methods.
- 15 You've, obviously, given a number of opinions in this case
- over a period of years; correct? 16
- 17 Given opinions in this case. I have provided written
- reports with opinions. 18
- 19 Do you hold all of the opinions contained in those reports
- with the same degree of certainty? 20
- 21 If I signed it, then I stand by it.
- 22 So the opinion that molecular weight markers is required,
- 23 for example, is held to the same degree of certainty as the
- 24 opinion that, for example, there is manipulation in a certain
- 25 figure; is that fair?

- I would say in my opinion, yes, molecular weight markers 1
- are required on gels. And as just mentioned, many, many 2
- journals agree with me on that. 3
- But at least one says it's prevalent not to have them; 4
- 5 correct?
- There is a publication in Plus One which criticizes the 6
- 7 field by saying that they are frequently left off published
- 8 images, but that publication does not say anything about raw
- 9 data.
- You know this is a criminal case; correct? 10
- 11 I am aware it's a criminal case, yes.
- 12 And it requires intentional misconduct; correct?
- 13 I am not aware of what standard is being applied in the
- 14 trial in order to reach a verdict --
- 15 Are you making an --
- -- intentional. 16
- 17 -- opinion in this case that there was intentional
- misconduct by Dr. Wang? 18
- 19 I have expressed an opinion that Dr. Wang committed
- 20 intentional misconduct, yes.
- Is it possible, in your mind, that Dr. Wang committed 21
- 22 negligent conduct that lead to at least some of the opinions
- 23 that you discuss in your reports?
- 24 Are we in trial already? I'm struggling to --
- 25 Q. It's a question.

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-- understand. We're in a Daubert hearing.
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   Α.
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Your Honor, I would ask him to answer 2 MS. BEIDEL: 3 the question.

THE COURT: Just answer the question. You are under 5 oath.

THE WITNESS: Can you ask it again, please? 6

BY MS. BEIDEL:

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- Is it possible, in your view, that Dr. Wang was negligent when he engaged in some of the conduct that you describe in the opinions in your reports?
- 11 Some of the conduct I would classify as negligent. 12 example, when we think about some of the Excel sheets that 13 appear to be some calculations in those sheets which could be 14 simply honest mistakes. Yes, there are. However, generally 15 speaking, in the field of scientific misconduct, there can be There can be two accidents. 16 an accident. There can be five 17 accidents. When there are hundreds of accidents, the -- and they all pile up, it becomes increasingly unlikely that that 18 19 can be attributed to incompetence or sloppiness.
- When you issued your opinions in the reports contained in 20 21 the exhibits, did you specify in which circumstance you were 22 talking about potentially unintentional error?
- 23 That is written in the reports. I believe in some of the 24 annotated Excel sheets you can see, for example, where there is 25 a formula that has been used to calculate the densitometry

- relative to the NR1 protein. And that formula has been copied 1
- Unfortunately, a dollar sign was inserted in the 2
- 3 formula and that has an effect of locking a particular cell.
- And so when that formula was copied and pasted instead of it 4
- 5 transposing down the sheet, everything got referenced to the
- wrong cell by mistake. That can be a common mistake from there 6
- 7 just being a dollar sign in the formula by accident. Yes.
- 8 that was described in the particular report.
- 9 So your intention is that if it was your opinion that
- 10 there could had been a harmless error caused for some issue you
- 11 would have indicated it?
- 12 Alternative explanations were described in the reports
- 13 regarding instances in which things could have come about by
- 14 error.
- 15 I want to talk about your method for a bit and focusing
- first on how it developed. So ultimately we see something that 16
- you called analytical pipeline for Western blot images. 17
- your method; correct? 18
- 19 That's a family of techniques that were used in this case,
- 20 yes.
- 21 And is that the method that you used to conduct your
- 22 analysis for purposes of this case?
- 23 Yes.
- 24 Have you used that method in other let's start with
- 25 publications?

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- A. Generally speaking, in the field of research integrity and scientific investigation, one does not publish one's findings in the scientific literature.
 - Q. So the answer is, no, you have not used those methods?
- Those methods have not been used in a manner in which the outcome has directly been published. The general reason for that is that when you're looking at published images in the scientific literature, the journals own the copyright of those images.
 - So, it's not possible, for example, to do what I have done on a number of the reports here which is to show a published image alongside an analysis of that image. A journal is unlikely to permit one of their figures to be reproduced for the purposes of pointing out that that figure has been inappropriately manipulated.
 - They're not likely to release copyright to allow that to occur, so, generally speaking, I'm not aware of anybody in this research integrity field who publishes their primary findings.
- Q. So, you are not aware of any paper that uses the three stage pipeline analytic method that you describe in this case; correct?
 - A. All of the methods described are in use in the research integrity field. They may come under different names, so essentially what is described in the report as a three-stage pipeline, a forwards analysis, that is something that is used

in the literature. A reverse analysis, that is something that 1 is used. Looking for unusual and additional features in 2

images, that is a method that's used. If they may not be 3

described in this exact way all synthesized in terminology, but 4

each of those methods has been used and while the findings of

those methods, the primary findings may not have been

7 published, there are numerous publications where people have

8 used these methods and made aggregate findings.

For example, regarding prevalence of particular types of image manipulation in the literature. Ferric Fang, Arturo Casadevall, Elizabeth Bik, there are a number of publications by these individuals where they have analyzed thousands of papers using these exact methods and then come up with broad conclusions regarding percentages of manipulated images in the literature.

- And you are certain of these exact methods?
- 17 They are a subset of these methods.
- A subset? 18

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19 I know for a fact that Elizabeth Bik, for example, uses 20 one of the tricks that is described in the report, very old 21 monitors, very old computer monitors, LCD monitors, not OLED or 22 modern laptop screens. The contrast of the image can be 23 adjusted by simply tilting the screen backwards, so one of the 24 techniques that Elizabeth Bik uses is for rapid screening of 25 images is open a paper, tilt the screen, open a paper, tilt the

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- screen as a quick way of looking to see if there are any 1
- undisclosed splicing seams within a Western blot because it's a 2
- 3 very quick way of looking at an image and seeing if it's
- 4 worthwhile for further analysis. That's one example of a
- 5 method --
- Did you use --6
- 7 -- that was described in the pipeline of adjusting
- brightness and contrast. That's a method that's been used to 8
- 9 publish. I don't use that method now because I have a new
- 10 laptop that has an OLED screen and that method no longer works.
- 11 MS. BEIDEL: Okay. Let's look at Exhibit-147,
- 12 please, Ms. Blackwood.
- 13 BY MS. BEIDEL:
- 14 You will see, Dr. Brookes, that this is an e-mail from you
- 15 to Special Agent Weeks dated October 6, 2022. Do you see that?
- Correct. Yes. 16
- 17 And it says that there are two files attached and one is
- an analysis outline. It says, analysis outline is the pipeline 18
- 19 methods document with edits as discussed.
- 20 Do you see that?
- Correct. 21
- 22 If we turn to the second page, this is at this point at
- 23 least the proposed analytical pipeline for Western blot images;
- 24 correct?
- 25 Α. This is, yes, a proposed pipeline.

- And what you mean by proposed pipeline is these are the 1
- steps you intend to take to conduct the analysis in this case? 2
- 3 Correct. Yes. Α.
- And so at this time at least you see on this page there is 4
- 5 three steps: Magnification, brightness and contrast adjustment
- and the curves function and Photoshop; correct? 6
- 7 Those are -- this is just the first page of the
- 8 document.
- 9 Right. And scrolling on, then there is recoloring and
- 10 PowerPoint, histogram analysis, error analysis tool called
- 11 Photo Forensics; correct?
- 12 Correct.
- 13 Finally, on the last page there is Image Twin and
- 14 sensitometry. Do you see that?
- 15 Yes.
- And nowhere in here is the droplet analysis, for example, 16
- 17 that you talked about; correct?
- 18 The droplet analysis is not listed in this initial
- 19 pipeline, correct.
- 20 And there was the example where you took data from the
- paper, and you took the graph and backed up to the Excel file 21
- 22 and analyzed that Excel file. That type of analysis is not
- 23 mentioned in this pipeline either; correct?
- 24 This is about analysis of images. This descriptive
- 25 document does not deal with how the images are obtained, so

- Page 121 of 206
- obtaining a PowerPoint and ungrouping and taking the images out 1
- of that PowerPoint, that is just the process by which the 2
- 3 images are obtained, but that is not described in this document
- which is about the analysis performed on the images once they 4
- 5 are in that.
- I'll rephrase that. My question was confusing. 6
- 7 talking about actually taking the graph that appeared in one of
- 8 the publication images where you were able to extract it and
- 9 get to the Excel spreadsheet that we saw with all of your
- 10 comment bubbles.
- 11 Do you know what I'm referring to?
- 12 The mechanism of extracting an Excel file from a
- 13 graph in a PowerPoint file was a later development during this
- 14 analytical process. I would argue that that doesn't come under
- 15 the agreement of image analysis, because what is being clicked
- on is not a Western blot image. What is being right clicked on 16
- 17 to extract that Excel file is a graph and a graph is not an
- 18 image.
- 19 Well, either way, at least at this phase, there was not a
- 20 step that said, for example, look at the graphs or look at the
- 21 data and see whether there is something corroborative of the
- 22 opinion that you could find in that?
- 23 That is described in the reports in which those
- 24 extractions were performed.
- 25 **Q**. Right. I'm focused on the method. In the method, there

is not a discussion of that type of step; correct? 1

- At the time this method document was written, the full 2
- scope of the type of data that would be made available to me to 3
- analyze was not clear so it would be unusual if not knowing 4
- 5 what file types I could expect to be given, I would
- predictively describe a method for extracting Excel sheets from 6
- 7 PowerPoint graphs, not even knowing the PowerPoint files
- 8 existed, and having been sent no PowerPoint files at that
- 9 stage, so --
- 10 So let's trace that --
- 11 -- as I said, this methods document is an early iteration
- 12 of a planned pipeline for image analysis, and as the process
- 13 continued, additional tools such as extracting Excel sheets
- 14 from PowerPoint files was added to the arsenal of methods that
- 15 were used.
- Let's trace the evolution of that analysis a bit. 16 Okay.
- 17 Let's show Exhibit-149. You'll see that this is now an e-mail
- from you to Special Agent Weeks dated November 14, 2022. 18
- little over a month after the last one. And it says, three 19
- 20 updated analysis pipeline describing three stages categories of
- 21 analysis.
- 22 Do you see that?
- 23 Correct.
- 24 And then if we move to the attachments, 149, you now see
- 25 what's called the analytic pipeline for Western blot images

- that has the three stages reverse analysis, forward analysis 1
- and further anomalies; correct? 2
- Correct. 3 Α.
- So that we have gone from the eight potential methods of 4
- 5 analysis down to this three stage reverse analysis, forward
- analysis and further anomalies? 6
- 7 That's a misconception. The -- this describes three
- 8 stages of the analysis. The previous document describes eight
- 9 tools to be used during these three stages.
- 10 And it's called analytic pipeline for Western blot images
- 11 here and we go to Page 2 of Exhibit-147, proposed analytical
- 12 pipeline for Western blot images.
- 13 Do you see that?
- 14 This is the early iteration.
- 15 Of the same thing; correct?
- This is an early iteration in which I called it a 16
- 17 pipeline. As you'll note, this does not have a number in the
- This was at a time where it was unclear how many 18
- 19 reports I would be asked to write and so, again, this is a very
- 20 early iteration in which I'm calling this a pipeline.
- 21 Subsequently, that nomenclature evolved to two documents, one
- being the description of the methods, and the other being the 22
- 23 pipeline for the application of those methods and those were
- 24 then numbered reports.
- 25 Let's go to Exhibit-87-4 at Page 15. I think you will see

that this is now or I'll represent to you that this is the 1

- current report as we understand it, report number three, with 2
- the three stages, reverse analysis, forward analysis and 3
- further anomalies. Do you see that? 4
- 5 Correct. And then on the first line you will see it says
- using the analytic tools described in document two. 6
- 7 Did you mean by that that you were going to use all eight
- 8 of those analytical tools?
- 9 It does not say that here.
- 10 So you mean --
- 11 It says, using the tools.
- 12 You mean you are going to do this three-stage analysis
- 13 pipeline using whichever of the tools you choose in that
- 14 particular circumstance?
- 15 Whichever are the most applicable for the data.
- example, an ORI droplet analysis, as you have seen, is more 16
- 17 appropriate for comparing bands in the white box image to the
- final published data. That would be less appropriate comparing 18
- 19 the white box image to the raw data because there are no bands
- in common to see if they overlay, so the tools depend on which 20
- 21 stage of the analysis we are and, indeed, on the individual
- 22 properties of the image.
- So, for example, in the case of the unusual square looking 23
- 24 black bands, in the extra image stage, that was an appropriate
- 25 place to apply a histogram analysis to show that those bands

- have different properties to the other bands in that document. 1
- You would not do a histogram analysis at every stage including 2
- the white box. 3
- So you use the tools that enable you to conclude there was 4
- 5 image manipulation?
- I used the tools that were most appropriate in order to 6
- 7 draw conclusions regarding the images.
- 8 If a tool would have, for example, indicated there was no
- 9 manipulation, you didn't use it; correct?
- 10 I frequently did. I frequently used ORI droplets.
- 11 frequently used histogram analysis. I applied tools because as
- 12 was already mentioned, we consider the number of alternative
- 13 explanations here for how these images came about. And so --
- 14 and as I've already described, as a scientist, it's important
- 15 to know when you are wrong and so you apply tools to -- to
- figure out if you are wrong. 16
- 17 But it's not as if you walked through each of the eight
- tools, you know, for every image you said number one, magnified 18
- 19 a pixel level detail. Number two, brightness and contrast.
- was not that regimented; correct? 20
- 21 There was not a regimented pattern for applying all eight
- 22 of the tools in all three stages.
- 23 Q. Okay.
- 24 For example, three times eight, that would be 24 separate
- 25 steps for every single image analysis. That would generate an

- inordinately large amount of paper. 1
- And the government's paying you \$150 an hour to do this 2
- work? 3
- The government is paying me a consulting fee for the 4
- 5 process of this analysis. That is in line with standard
- consulting fees for this type of work. 6
- 7 But eight steps for each image would have taken longer and
- 8 cost more; correct?
- 9 Doing 24 separate steps would have drastically increased
- the amount of time and, therefore, billable hours to the 10
- 11 government.
- 12 I don't want to take you through with every single one of
- 13 the reports. I'm conscious of your time and the Court's time.
- 14 But let's just talk about a couple of examples.
- 15 Can we go to Exhibit-101A? I'll represent to you, Dr.
- Brookes, that this is a red line run between different versions 16
- 17 of your reports that were produced to us in discovery with the
- newest showing up as the newest. It's a red line. 18
- 19 Do you see that?
- 20 Α. Yes.
- If we turn to Page 13. This, by the way, is the standards 21
- 22 and expectations section talking about Western blotting in
- 23 general.
- 24 On page 13, we will see there is a section that was
- 25 removed called shared noise features between image layers. Do

you see that? 1

- 2 Α. Yes.
- 3 And toward the end it says, quantitation of the degree of
- overlap between any two images is difficult by using image 4
- 5 enhancement techniques as described here. It can often be
- shown that two images likely arose from a single origin if they 6
- 7 share more features than we expected by --
- 8 Where exactly --
- 9 THE COURT: Let the attorney restate the question so
- 10 the court reporter can get it, at least the part you were
- 11 reading.
- 12 BY MS. BEIDEL:
- 13 While quantitation of the degree of overlap between any
- 14 two images is difficult, by using image enhancement techniques
- 15 as described herein, it can often be shown that two images
- likely arose from a single origin and then it continues on. 16
- 17 Do you see that?
- I see where that is written. 18 Yeah.
- 19 101A?
- 20 In an early draft of that document.
- 21 Why did you remove that particular sentence?
- 22 I believe upon consultation with Agent Weeks and Mr.
- 23 Tyler, it was thought it would be a good idea to shorten this
- 24 document, and so this entire section was removed, including the
- 25 text above and below it. So it was not a choice necessarily to

remove just that particular sentence. It was decided to just 1

- not to have a recap section so that entire thing, it was 2
- 3 thought that did not add anything to the document.
- Do you still agree with your statement that quantitation 4
- 5 of the degree of overlap between any two images is difficult?
- Quantitation of the degree of overlap between two 6
- 7 images is difficult. Just because something is difficult does
- 8 not mean it's impossible.
- 9 Do you still agree that it can often be shown that two
- 10 images arose from a single origin, but not always. Do you
- 11 still agree with that?
- 12 Correct. Often, but not always. Very, very few things
- 13 are always true.
- 14 For example, I've seen you reference as a potential,
- 15 quote, smoking gun that if you say duplication of two bands or
- two blots within a paper that would be a smoking gun evidence 16
- 17 of manipulation; is that right?
- I believe you are referring to an e-mail in which the 18
- 19 phrase smoking gun was used and the specific criteria that was
- 20 mentioned there is whether the noise pattern in two white box
- 21 images was reproduced. That would be a smoking gun because for
- 22 the noise pattern and in the white box images to be reproduced
- 23 would be highly, highly unusual. There are, in fact, two
- instances -- three -- sorry. Report number 4.5, report number 24
- 25 6.4 and, in fact, the original data set that I was sent

regarding the problem with the original data provided to the 1

- Journal of Prevention of Alzheimer's Disease. 2 Sorry. The
- 3 Journal of Alzheimer's Disease and Therapy. The original data.
- 4 In all those cases, there was, if we are going to call it a
- 5 smoking qun, there was -- that criteria was met which is the
- reproduction of patterns of noise across multiple white box 6
- 7 images which is physically impossible.
- 8 And you're basing that on the background noise in the
- 9 image; correct?
- 10 Correct. The background noise of two independently
- 11 generated images cannot be identifiable.
- 12 Let's look at Exhibit-106. This is your report 4.5 which
- 13 I believe is one of the reports you just mentioned?
- 14 Correct.
- 15 If we turn to Page 5. Are you saying that the background
- noise is repeated within the same white box and that is what is 16
- 17 impossible?
- In this particular instance, yes. That is noise that is 18
- 19 fuzz that is, you know, static on the television screen from
- 20 the old days, that is random noise. That noise should not be
- 21 reproduced in multiple places across an image. One possibility
- 22 is digital cloning of an area of an image which would lead to
- 23 such reproduction of the noise pattern.
- So what about this versus this? 24
- 25 THE COURT: I'm sorry. Can you speak into the

1 microphone, please?

BY MS. BEIDEL:

- 3 Q. What about the -- I'm drawing two circles on the images.
- 4 One on the top portion of noise and one on the bottom. They
- 5 don't look the same to me. Do they look the same to you?
- 6 A. Yes. There are numerous shad features between those
- 7 areas.

- 8 Q. You see the same darkened line at the bottom of the top
- 9 | line as you do the other. Do you see that in both?
- 10 **A.** The darkened line is different because there is an ever so
- 11 slightly different one pixel shift in where the box was drawn.
- 12 There are numerous other shad features. For example, if we
- 13 look, there are three little dots here. Those same three
- 14 | little dots appear up here. There are lots of -- there is
- 15 another triangle of dots that appears here. That same triangle
- 16 of dots appears here. We can look anywhere in this image and
- 17 argue that the pixel level, the simple thing to do would be an
- 18 ORI droplet analysis to overlay these images. I would be more
- 19 than happy to do that, if necessary, but I think to the naked
- 20 eye, you can see that the noise pattern here is the same.
- 21 Q. So, for example --
- 22 **A.** And that is just highly, highly unusual within -- within
- 23 | an image like this.
- 24 | Q. Choosing another portion of the image at random, I have
- 25 just drawn a circle. Here is the other image. I see three

- bigger dots in the top one. I don't see them in the bottom. 1
- Do you see that difference? 2
- There is a slight difference in the intensity of those 3
- dots, but the features above and below are the same. 4
- 5 So, when the differences favor your opinion, you say they
- amount to manipulation, but when they don't, they are 6
- 7 explainable; is that right?
- 8 That's not correct. That's not what I'm saying.
- 9 Are you saying that an ORI droplet would be a better
- 10 indication of this difference than this noise analysis?
- 11 And I apologize for not including an ORI droplet in
- 12 this report.
- 13 But, yet, the noise analysis was a smoking gun a second
- 14 ago?
- 15 I believe the phrase "smoking gun" was used in a different
- time to when this report -- I would have to look at the time 16
- 17 line of when this report was used. Sorry. When this report
- was written and submitted. 18
- 19 If ORI droplet analysis is the preferred technique, why
- isn't your method just simply we did an ORI droplet analysis? 20
- 21 There are several steps needed to get to that point.
- 22 many cases, as seen here to the naked eye, it is often not
- 23 necessary to do the ORI droplet analysis. You can see straight
- 24 away that the image noise is the same or the bands are the
- 25 The ORI droplet analysis is essentially a confirmatory same.

- technique to confirm something that is very often quite obvious 1
- 2 already.
- 3 There is lots of techniques either that have been
- 4 developed or being developed that use technology. You've
- 5 mentioned Photo Forensics and Image Twin as two examples;
- right? 6
- 7 Image Twin is one that I use quite regularly, yes.
- 8 There is others out there that are also available?
- 9 There is another one that was developed by Edward Delp at
- 10 Perdue University in Indiana. That is I believe called Sila,
- 11 S-i-l-a. I actually provided some of the original data to
- 12 Dr. Delp for the development of that algorithm.
- 13 Are you aware of studies that test the error rate of any
- 14 of those algorithms? Sila or Image Twin or Photo Forensic or
- 15 any others?
- 16 I believe is Sila one, as part of their reporting, and the
- 17 original paper, they conducted a false positive, false negative
- and an error analysis in that published paper. I would have to 18
- 19 go back and read the published paper. The authors or the
- 20 creditors of Image Twin, I believe there have been a couple of
- 21 third-party reports on that in the published literature.
- 22 However, that is a proprietary algorithm, so I'm not aware that
- 23 the authors of Image Twin have published anything about its
- effectiveness or how it works behind the scenes. 24
- 25 **Q**. Okay. And you didn't use Sila as part of your analysis;

correct? 1

- No, I didn't. Shortly after the publication of that 2
- 3 paper, the website for the download, the get-up site, the Sila
- software is essentially no longer available. For whatever 4
- 5 reason, Dr. Delp chose to pull the API. I believe he's in the
- process of making that software available on a commercial basis 6
- 7 to journals so, therefore, probably pulled the API from getting
- 8 up in order to prevent it from being freely distributed.
- 9 Image Twin you did use, but you are not aware of whether
- 10 there is data on its accuracy because it's proprietary?
- 11 I have used Image Twin. However, specifically I have not
- 12 used Image Twin in this case.
- 13 Oh, you didn't. Okay.
- 14 The reason for that is because like a lot of AI-type
- 15 applications, there are privacy concerns. The files that were
- provided to me were on a confidential basis by the government. 16
- 17 It is unclear to me when loading things in to an AI, whether
- those images will be used in training, for example, by that AI. 18
- 19 And so none of the images that I was provided by the government
- were sent to any third-party software AI-type platform for 20
- 21 confidentiality reasons.
- 22 Is Photo Forensic an AI tool?
- That is a simple analysis which takes place on the 23
- 24 It runs within the browser. desktop.
- 25 Q. So you did use that tool because those AI-type concerns

1 don't apply?

- 2 A. At the time -- in fact, I think Image Twin was launched in
- 3 | mid-2023. I was a beta tester for it. But Image Twin and
- 4 other AI tools are very, very recent developments in this
- 5 | field. At the time, when I was doing most of this analysis in
- 6 late 2022 and early 2023, those tools were not yet available.
- 7 Q. But there are, as we sit here today, AI-based tools
- 8 available that could analyze Dr. Wang's image and as far as you
- 9 know, they were not used in this case?
- 10 **A.** As far as I know, they were not used in this case.
- 11 **Q.** Just one second. Apologies. I want to look at a few more
- 12 of your individual reports. Just a second here. Let's go to
- 13 Exhibit-130. You'll see this is one titled, origin of Western
- 14 | blot images and figures from a particular N.I.H. grant
- 15 proposal. Do you see that?
- 16 A. That's correct. This is an unnumbered report, so I
- 17 | believe this may have been a very early example. In fact, you
- 18 can see the opening paragraph of this describes the introduct
- 19 | rather than report number two as it would otherwise be referred
- 20 to.
- 21 **Q.** So, let's look at Page 2 for a second. Just before those
- 22 | two boxes. It says, the images appear to originate from x-ray
- 23 | film. Do you see that?
- 24 A. Correct.
- 25 \mathbf{Q} . You use that appear to language pretty frequently

- throughout your reports. Appear to originate from some source. 1
- Do you recall that? 2
- 3 Correct. Yeah. Α.
- Why do you say that appear to originate as opposed to just 4
- 5 concluding that they do originate from the x-ray film?
- I base my description on what appears. 6
- 7 Okay. You are just doing a visual analysis essentially by
- 8 some tools of what appears; correct?
- 9 At this stage, this is a visual analysis saying what these
- 10 images appear to be. You are holding a piece of film in your
- 11 I think you can agree the size and dimensions, the fold
- 12 in the corner, the curved edges -- right -- these are all
- 13 common properties of a piece of x-ray film. So that is a
- 14 reasonable appearance for a piece of x-ray film.
- 15 MS. BEIDEL: Let's blow up the two pieces of film at
- the top a bit, Ms. Blackwood. Okay. 16
- 17 BY MS. BEIDEL:
- So, using this just as a representative example, assume 18
- 19 with me for purposes of this discussion that that white box is
- 20 pasted on top of what appears on the left. Okay.
- Correct. 21
- 22 Then couldn't that white box have derived from some other
- 23 film that accurately reflects the data that ties in to the
- 24 N.I.H. grant?
- 25 As we have already discussed, that could be another image

- that contains the bands in the white box image.
- And if you are doing analyses, for example, of these top 2
- two very dark blots to prove they're identical, that only 3
- proves that the background on which the white box was pasted 4
- 5 matches the image on the left; correct?
- So, when you say the white boxes are pasted, are you 6
- 7 referring to the white box or the white box and everything in
- 8 it, or the white box and the bands in the white box, but not
- 9 the noise in the white box? Is that what you are referring to
- 10 when you say it was copied and pasted?
- 11 What I meant to refer to is the entire white box and all
- 12 of its content including the noise and the bands.
- 13 It is not possible that the entire white box could had
- 14 been pasted in. If that were to be the case, it would obscure
- 15 the noise from the original raw image and yet what we see here
- is a scene in many of the reports: The noise in the raw image 16
- 17 appears again in the white box image.
- So simply pasting those bands on top is impossible to see 18
- how the underlying noise would come through to show up in the 19
- 20 final image.
- Couldn't the white box have derived from a different 21
- 22 exposure of those same two?
- As we have already mentioned, different exposures show up 23
- 24 on different pieces of film. These are the same piece of film.
- 25 **Q**. Not if the white box is pasted on top. It could have come

- from a different piece of film in that scenario? 1
- The white box could have come from another piece of film, 2
- but then it would bring with it the noise from that piece of 3
- film and would not show the noise from the raw piece of image 4
- 5 on the left.
- I'm going to switch the document camera for a second. I'm 6
- 7 showing the film that is contained within Exhibit-52.
- 8 believe what you told me is these are all different exposure
- 9 times of the same two pieces of film; is that right?
- 10 Correct. So, if you move the document down a little bit,
- 11 you can see in the top corner it says 60 seconds. On the other
- 12 side it says 30 seconds written backwards, so the film was
- 13 flipped to collect those different exposure times.
- 14 So the one on the right was exposed for 30 seconds and the
- 15 one on the left was exposed for 60 seconds, but it's otherwise
- 16 the same?
- 17 Correct. Correct.
- And you see this dot right here? 18
- 19 Correct.
- 20 And it appears over here; correct?
- 21 Correct.
- 22 And this could just as easily been two separate pieces of
- film as it is one film with two blots? 23
- 24 It could be two separate pieces of film. However,
- 25 positioning that film some people do sometimes cut the film up

into numerous pieces. So positioning the film in exactly the 1

- same place in the cassette when you have numerous little, small 2
- pieces of film, is -- is very very difficult to do properly. 3
- Assume we did have two little pieces of film, that 4
- 5 background noise would match, even though these are different
- exposures, wouldn't it? 6
- 7 It would in terms of these being different exposures,
- 8 correct.
- 9 So, if there was a --
- 10 What I have done is to collect these exposures, and so the
- 11 raw data is self-contained within one of those pieces of film.
- 12 There would be absolutely no scientific reason for me to have
- 13 one exposure of a piece of film and then to take the set of
- 14 bands and to transpose that set of bands on top of another
- 15 piece of film containing noise and then to merge those images
- in order to maintain some of the noise, but some of the bands 16
- 17 but different bits of each image. That is not how Western
- That also still does not answer the issue of 18 blots are done.
- 19 where is the raw data and the original image from which the
- 20 bands in the white box appeared.
- But it's possible for noise to match between two different 21
- 22 exposures; correct?
- 23 It's possible for noise to match between two different
- 24 exposures. It is not possible, as we discussed, for noise to
- 25 match between, say, stripping and reprobing. But different

- exposure of the same blot within a few seconds of each over, I 1
- put a piece of film down, take an exposure, flip it around, 2
- 3 take a second exposure, that can sometimes lead to the same
- noise pattern. 4
- 5 So, if the white box derived from a different exposure of
- the two blots on the bottom of the left image and then those 6
- 7 were pasted on top of that image, isn't it possible that the
- 8 noise remnants came from that different exposure?
- 9 It would be highly unusual because what you see here is
- 10 the white box image has a very very clean background.
- 11 white box image is brighter than the film behind it, so the
- 12 film behind it is sort of a muted gray. Just all of this stuff
- 13 is gray.
- 14 So the white box image being much brighter. When an image
- 15 is brightened, the noise should become less prominent.
- would be very unusual for the white box to come over in a much 16
- 17 brighter format with the bands and yet the noise comes over
- darker. 18
- 19 The most logical explanation here is that the noise
- 20 originated from the raw image on the left because the noise in
- 21 that image is much darker and more prominent to begin with, so
- 22 pasting a very light-colored image on top of a dark-colored
- 23 image, the dark image is going to be the prominent source of
- 24 noise in that image.
- 25 When you say the most likely --

- 1 A. Again, it is infeasible scientifically as to why anybody
- 2 | would choose to prepare an image in this way.
- 3 **Q.** When you say the most likely explanation, you mean
- 4 51 percent?
- 5 **A.** No. I mean, the likely explanation for the appearance of
- 6 the noise in the white box image is that this noise comes from
- 7 taking the raw image on the right and brightening a particular
- 8 | section of that image resulting in the appearance of the light
- 9 noise patterns which are seen in the white box image, for
- 10 example, this here.
- 11 Q. But it's possible that it came from a different exposure,
- 12 just not likely in your explanation?
- 13 **A.** It is possible, but highly, highly unlikely that
- 14 the bands here and the informational content of the white box
- 15 | image, it is possible that that came from another unknown
- 16 | source. We were not provided with -- with those images if they
- 17 exist. It is also, it should be reiterated when thinking about
- 18 the reverse and the scientific chain of provenance and custody,
- 19 that other exposure, if it exists, is those are the bands that
- 20 appear in the final paper. That other exposure should had been
- 21 | the piece of film that was retained. That is the raw data for
- 22 that image and yet, that raw data does not appear to exist.
- 23 **Q.** Do you --
- 24 | A. What we have instead is another piece of film which
- 25 appears to share some noise, but does not contain the bands.

- Do you know how Special Agent Weeks obtained these films 1
- that he provided to you or blots that he provided to you? 2
- I don't know. These are digital images. They're JPEGs. 3
- My understanding is that Agent Weeks obtained them as JPEGs. 4
- 5 You don't know whether he, for example, executed a search
- warrant at CUNY and obtained everything available in Dr. Wang's 6
- 7 lab?
- 8 I don't know. That's beyond my involvement in this case.
- 9 There could easily be other records, other devices, other
- 10 digital images available in that lab and you would not be aware
- 11 of it; correct?
- 12 There may be. I'm not aware if such resources exist.
- 13 You do know that CUNY was conducting its own internal
- 14 investigation; correct?
- 15 Yes. I already mentioned that, yes.
- Do you know whether they obtained files from Dr. Wang's 16
- 17 lab at any point?
- I would say that it is typical during 18
- investigations for institutions to request the files and other 19
- materials from laboratories. 20
- 21 Do you know whether it's possible that CUNY took some of
- 22 those files and never returned them in a way that they were
- 23 available to Special Agent Weeks?
- 24 I don't know the timeline in which any files were taken or
- 25 obtained so, again, this -- that is beyond my involvement in

- All I can do is base my analysis upon the materials 1 this case.
- that I was provided with. 2
- What you are doing, though, is saying that, in your view, 3
- it would be standard scientific practice for Dr. Wang to have 4
- 5 retained these other films, if they exist; correct?
- Correct. 6
- 7 Isn't it true that you don't know one way or another, as
- 8 you sit here today, whether Dr. Wang retained those films?
- 9 Α. Correct.
- 10 So isn't your entire opinion based on the premise that
- 11 there are no other films so that there is image manipulation
- 12 based on the images that you saw only?
- Can you repeat the question? 13
- 14 I can, and hopefully I will do it better this time.
- 15 Isn't your entire opinion based on the comparison of the
- images that Special Agent Weeks provided you with only and 16
- 17 completely to the exclusion of any other images, if they exist?
- My opinions and conclusions are based on the images that I 18
- 19 was provided with. What I can conclude, for example, is that
- 20 in numerous cases the bands pattern in the white box image and
- 21 the final published image does not find any provenance in the
- 22 raw image. That is the extent of my conclusions.
- 23 Let's go back to Exhibit-130, which we were looking at a
- 24 second ago, Page 18?
- 25 THE COURT: So, Ms. Beidel, it's now one o'clock.

think we are going to have to take a lunch break. 1

I'm almost finished, I think, Your 2 MS. BEIDEL:

3 I was going to ask like two more questions and then

consult with my client to see if he had anything else and I can 4

5 wrap up if that works for you.

With the caveat that, I've never had a 6

7 lawyer say two more or one more question and actually stick to

8 it, we can go two more minutes.

> MS. BEIDEL: Thank you, Your Honor.

BY MS. BEIDEL: 10

- 11 You will see in the last paragraph here it starts with, in
- 12 the absence of exculpatory evidence, e.g., original blot films
- 13 with lower exposure that, indeed, show that the ends of
- 14 interest at 90 kilodaltons and 280 kilodaltons as they appear
- 15 in the proposal figures, it is my professional opinion that the
- figures as presented in the proposal are fabrications. 16
- 17 Do you see that?
- 18 Yes.
- 19 Do you still agree with that as you sit here today?
- 20 I agree with that opinion as expressed in this
- document. 21
- 22 So, if there were exculpatory evidence, e.g., original
- 23 blot films with a lower exposure that showed the bands, then
- you would change your opinion; correct? 24
- 25 Α. I would change my opinion if such evidence became

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available. 1

- MS. BEIDEL: One second, Your Honor. 2
- We can take that down, Ms. Blackwood. 3
- One question from Dr. Wang and I'm going to stick to my 4
- 5 deal with you.
- BY MS. BEIDEL: 6
- 7 We have seen film example that's a whole 8-by-11 sheet;
- 8 correct?
- 9 Yes.
- Is it possible to use films for Western blots that are 10
- 11 smaller, you know, a strip that shows two blots, for example?
- 12 It can be common to cut up pieces of film in to smaller
- 13 sizes. There is a limit to how small those can get because the
- 14 film eventually has to be fed into a developer machine. And if
- 15 the film is too small, there is quite literally a risk that it
- get lost inside the machine because it's too small for the 16
- 17 rollers inside the machine to handle. So there is a lower
- limit of roughly a third of a piece of film. Any smaller than 18
- 19 that, you can't feed the film in to the developer machine.
- 20 MS. BEIDEL: One more second.
- BY MS. BEIDEL: 21
- 22 If you are using small strips of film, could they be taped
- on top of a larger piece of film for purposes of development in 23
- that process you just discussed? 24
- 25 Α. No. You would break the developer machine. The developer

Case 8:24-cr-00211-TDC

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machine can only handle one piece of film at a time.
 1
    a common mistake that's made is the box of film comes with a
 2
    piece of cardboard, the first piece, and it's a common rookie
 3
    mistake to accidentally feed the piece of cardboard into the
 4
 5
    machine. If you feed anything other than one piece of film at
    a time into the Western blot developer, you will break the
 6
 7
    developer machine and end up with a $30,000 bill from your
 8
    department chair. So that's not good.
 9
         Well, that's not good. There could be a strip that's
10
    maybe a third of a sheet is a fair size?
11
         There could be a piece of film which is a third of a piece
12
    of film. However, as you see in many cases, many of the white
13
    box areas were two by three centimeters which is simply way too
14
    small to feed into any type of developer machine without
15
    breaking the machine.
              MS. BEIDEL: Nothing further, Your Honor.
16
17
              THE COURT:
                          So, Mr. Tyler, where are we now? Do we
18
    have any redirect?
19
                          I think just one point I would like to
              MR. TYLER:
20
    clean up. Is that acceptable?
21
              THE COURT:
                          You are testing the patience of -- not
22
    just me, but staff. We have been at this -- it's a lunch hour.
23
    So how many questions?
24
              MR. TYLER: Just a few questions just to try to clean
```

25

up one thing.

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THE COURT: Okay. Go ahead.

2 MR. TYLER: Ms. Cornich, could you pull up

3 Exhibit-5A?

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4

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25

REDIRECT EXAMINATION

BY MR. TYLER:

Q. Dr. Brookes, when Ms. Beidel was asking you about assumed images and other images that may exist, during our direct examination this is one image. If we go to Page 6 here, please. We matched up the boxes here on this particular image and I think with respect to any additional images, would those have to be digital images or film image based on the analysis

- 12 that you did here?
- 13 **A.** The superimposed images would have to be digital because
 14 the noise is digital. The noise between the raw and the white
 15 box image is digital, so anything that is pasted in here has to
 16 be digital.
- 17 MR. TYLER: Okay. No further questions, Your Honor.
- 18 **THE COURT:** Okay. Anything else just on that?
- 19 MS. BEIDEL: No, Your Honor. Thank you.
- THE COURT: Okay. Could I just ask one question,
 possibly more. Just to clarify, since we talked about this
 noise analysis of what is in the background of one image versus
 another. Is there any known or any established standard on how
 many or how many points of comparison between one image and

another that you would need to see to determine whether the

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background noise means that they are the same or that they are
 1
          Is there any established industry, but sort of scientific
 2
    standard has been adopted across the community?
 3
                            Insofar as this is a field, it's a very
              THE WITNESS:
 4
 5
    young developing field, the standards have not yet really been
 6
    defined.
              There are some standards from other types of image
 7
    analysis in the scientific field. For example, with
 8
    fluorescent microscopy you may have an image of some cells and
 9
    you may have two proteins and you want to show whether those
    two proteins overlap on a microscope, that can be done with
10
11
    very sophisticated software which most people do not have
12
    access to which is essentially provided by the people that sell
13
    the microscope.
14
         So, if you have a Leica microscope, that's about a half a
15
    million dollar microscope, that comes with a proprietary
    software package that will enable you to line up images and say
16
17
    this image and this image overlap by 94.3 percent and put a
    number on a pixel basis. That is not software that is commonly
18
19
                That can be done, but my lab typically does not use
    available.
20
    fluorescent microscopy. I know of only one other lab in my
21
    university that has access to that type of equipment and
22
    software.
23
         So it's not something that, for example, Elizabeth Bik or
    other people operating in this area would have access to.
24
```

Second question on this. ORI, do they

25

THE COURT:

have any tools that help, are available to do that kind of 1 comparison? I know they have the droplets, but the droplets 2 3 seem to be for something else. THE WITNESS: Yeah. The ORI, as far as I know, does 4 5 not make available a quantitation tool such as that. THE COURT: Thank you. Anybody want to follow 6 Okay. 7 up on that they can. 8 MR. TYLER: Nothing from me, Your Honor. 9 MS. BEIDEL: No, Your Honor. Thank you. 10 THE COURT: Thank you very much, Dr. Brookes. 11 appreciate you testifying. I know it was a long morning in to 12 the afternoon. So you can step out now. 13 (Witness excused.) 14 15 THE COURT: So what we are going to need to do from a scheduling standpoint is we will take a lunch break. 16 we have a -- let me just check. We have a sentencing at 2:30, 17 but I would like to hear from counsel on these issues after 18 19 lunch and I don't think Dr. Brookes needs to be here unless 20 anyone thinks we might need him again. I know he has a 21 conflict at some point in the afternoon. We could perhaps come

back -- again, I want to make sure the staff gets a sufficient

break, but I'm willing to come back at let's say ten minutes to
two and see what we can accomplish before the 2:30 sentencing
and let's see where we are.

22

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Is that okay with everyone?
 1
                          It is for the government, Your Honor.
 2
              MR. TYLER:
              MS. BEIDEL: Yes, Your Honor.
 3
              THE COURT:
                          I take it there is no further questions
 4
 5
    at this point, even though I think I had mentioned indirectly
    that the defense experts were still in the picture but,
 6
 7
    obviously, there is some connection between what happens to Dr.
 8
    Brookes' testimony and them, so we are starting here. So I'll
 9
    see you at 1:50.
10
              MS. BEIDEL:
                           Thank you, Your Honor.
11
              DEPUTY CLERK:
                             All rise. This Honorable Court now
12
    stands in recess.
13
         (Whereupon, a lunch recess was taken from 1:11 until 1:54
14
    p.m.)
15
              DEPUTY CLERK:
                             All rise.
                                         This Honorable Court
16
    resumes in session. The Honorable Theodore D. Chuang
17
    presiding.
              THE COURT:
                          Thank you, everyone. Please be seated.
18
19
    So my thought was that on the one hand the defense has filed a
20
    motion to exclude Dr. Brookes, but I think I might want to
21
    start by asking questions of the government just to understand
22
    the scope of the opinions that you are actually planning to
    offer at trial.
23
24
         As I read it, there is a lot of reports from Dr. Brookes,
25
    and now I am finding out from the testimony that some of them
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might have been sort of drafts or earlier versions, and I know there is some I never even got, which is fine up until now.

I want to get a handle on what exactly the testimony would be and then we can see who is in the best position to argue about it from there.

So, Mr. Tyler, maybe you can give me -- usually like in an easy situation you just have like one expert report that lists like five things, and that's what we are going to hear, no more and no less.

Can you distill for us now what you expect him to testify about?

MR. TYLER: Yeah. So we -- I think also was probably something the defense raised, Ms. Beidel raised earlier, is we don't expect any of his testimony to sort of like be outside of the bounds of the exhibits that were admitted today.

Now, there are a lot of reports in there that we didn't have time to go over, but a lot of them are repetitive in the sense that there is like the reverse analysis, the forward analysis, and like that is very similar to what we did in court today. And it is just for a cross -- I think it's like 13 or 14 unique images across the entire case like some of them are viewed across multiple grants and multiple papers.

But the reason that we, frankly, did some of those other vignettes afterwards was to show some of the other pieces that are sort of in that third part of his analysis where he found

something else out.

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So I think we covered substantively -- I can't say this absolutely, but I think we covered almost all substantively of what the different types of analysis are that are sprinkled through the reports with like, for example, reverse analysis would be repeated in basically all 13 of the images, and the same with the forward analysis. And then the extra pieces of analysis just depends on whether it was available for that particular image or not.

THE COURT: So how many different images are there? Thirteen?

MR. TYLER: I don't have it pulled up right in front of me, but that is right around there, yes, Your Honor.

Going in to trial, it would be helpful THE COURT: for me to understand -- and we don't have to do this today, but even though it might be in these binders, it would be helpful for me to know if he's going to testify about 13 different images, are there 13 different reports and just looking at the final versions, at least that's what I would start with. understand maybe there is some arguments to be made if things have changed over time. Are there four images that are covered in a single report? It would be good to have sort of what we think is the expert report regarding those 13 images, whether it's 13 different reports or some smaller number.

> MR. TYLER: Yeah. It's going to be -- it's going to

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be 13 -- it's going to be at least 13. Each report
 1
 2
    essentially -- actually it might be more than 13 because some
 3
    of the images --
              THE COURT:
                          Thirteen images or 13 reports or both?
 4
 5
              MR. TYLER:
                          Thirteen -- it's going to be 13 images.
    It's going to be more than that number of reports. I don't
 6
 7
    have that right in front of me.
 8
         One thing that Your Honor might -- what might be helpful,
 9
    you may recall back in I think December we filed I think under
10
    seal a summary of all the different images in each of the
11
            That is -- those are the same figures with one
                There is one additional figure that we notified the
12
13
    defense about, but that is -- those are all the figures in each
14
    of the grants that we're going to cover.
15
                         Do you have the ECF number for that?
              THE COURT:
16
    was kind of a while ago.
17
              MR. TYLER:
                          Yeah. Exhibit-68. It's the exhibit to
    ECF 68.
18
19
                          We will go back and look at that.
              THE COURT:
         So there is these 13 different documents. At some point,
20
21
    by the way, it would be helpful to get a little chart with like
22
    which report goes with which and maybe even getting all the
23
    reports together because the reports probably aren't really
24
    evidence, but they are going to be useful for everyone.
25
    Probably the subject of cross-examination, if nothing else.
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So, for each of those, there is at least the potential for
Dr. Brookes to testify about how there is three pieces, the raw
data slide, the white box thing and then the published version.
          MR. TYLER:
                      Correct.
          THE COURT: And he wants to be able to say that --
well, actually can you just clarify for me the white box thing?
I'm not sure -- I was a little bit surprised by Dr. Brookes'
answer. It sounds as if nobody does white boxes. This is the
only time that's ever happened.
     Can you just explain what your understanding of the white
boxes are?
          MR. TYLER:
                      Yes.
          THE COURT:
                      The defense may have a different view,
but go ahead.
                      I imagine defense does have a different
          MR. TYLER:
view.
     Our understanding is that like if you are going to do
image analysis, I think Dr. Brookes was trying to describe, is
like if you are going to do that, you do that across the entire
blot, and so seeing like these little pieces is not
something -- it's not something we have seen in the people that
we have talked to as a common practice for any of the people.
          THE COURT:
                      But none of them were considered final
images; right?
          MR. TYLER:
                      That -- I mean --
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THE COURT: What we call the white box document? So usually they are like -- if you look MR. TYLER: at the white box, that is usually directly transferable to the final image. Like it appears that that is like essentially like cropped and then put into the final image. THE COURT: So one opinion is that those are the Another is that the white box and the so-called raw document derive from the same place. Less from the blots, but from this background noise concept. And the third piece is that the blots in the white box can't really be replicated or created based on the raw image. MR. TYLER: Yes. That is all -- all true. third -- the one caveat is the third scenario, what he calls his third step is really some of the other stuff we covered where, in other words, where there is 12 bands instead of -his third step of the analysis is there is some other feature that he's noticed that like is usually specific to that particular analysis, like there is 12 bands instead of 13 and there is like -- there is like multiple images that correspond to the same raw image. So those -- I do think we hit most of those today in the testimony, so those pieces are going to be a little bit more --THE COURT: Do those come up multiple times? just gave me one example, or do they only come up once each? MR. TYLER: Most of those only come up once each.

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THE COURT:
                          One was, for example, the 12 and 13.
 1
    was sort of the shape of the --
 2
                                  The 12 and 13 only comes up once.
 3
              MR. TYLER:
                          Yeah.
                          The shape of the band?
              THE COURT:
 4
 5
                                 The shape of the band --
              MR. TYLER:
                          Yes.
                          What about this pH scale issue?
              THE COURT:
 6
 7
              MR. TYLER:
                          That also only comes up once.
 8
              THE COURT:
                          And then what about this terminal digit
 9
    argument?
10
              MR. TYLER:
                          Yeah.
                                  That also only comes up -- well,
11
    there might be a couple other -- he makes the same analysis on
12
    different sets of data, but it's the same -- it's the same.
13
    You take data inputs and then you do the statistical.
14
                          But are you going to have him testify
              THE COURT:
15
    about data that is not directly associated with one of these
            Are we going off in another area, or is it just data
16
    that connects to the images?
17
                          So this -- so this is like a little -- so
18
              MR. TYLER:
19
    simply the one he talked about today was tied to a particular
20
    journal article. I don't remember -- there is a number of
21
    figures in that article that are also repeated N.I.H. grants,
22
    but I do not think all of them are.
         So that is -- that is -- I'm just trying to be totally
23
24
    transparent about my understanding of that.
25
              THE COURT:
                          When you say "figures," you are talking
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about the images, not the data numbers?
 1
 2
              MR. TYLER:
                          Sorry, Your Honor. In other words, if
 3
    you have a journal article with 13 figures in it, maybe six of
    them are ones that are also like essentially also appear in a
 4
 5
    grant application, and that terminal digit that houses was
    against all 13 in the actual -- in that actual article.
 6
 7
              THE COURT:
                         Okay.
                                 Then when he's talking about data,
 8
    whether it's relating to the terminal digits or the pH scale, I
 9
    was not at all aware of this issue about pulling up data from
10
    the PowerPoint presentation -- PowerPoint I guess if you -- is
11
    this how this works? Is that you have a spreadsheet and you
12
    ask PowerPoint to make a graph basically?
              MR. TYLER:
13
                          Yes.
                          So it's the underlying data?
14
              THE COURT:
15
              MR. TYLER:
                          Correct.
16
              THE COURT:
                          Is all the data in that form, or is some
17
    of it just in some other freestanding set of data?
                          So, I mean, it's in both. Right.
18
              MR. TYLER:
19
    Because the PowerPoint -- basically the PowerPoint has the
20
    chart embedded in it. Within that chart, what we tried to show
21
    was that you can basically unpack the Excel that is the input
22
    for that chart, and so that -- like that was that extraction I
23
    tried to show. We can think more about how we can present that
24
    at trial. Frankly, some of it could probably be done through
25
    testimony, but that's -- the idea is that the PowerPoint has a
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chart in it and within that chart is embedded data that is the
 1
 2
    source for the chart.
                          And that's the only data we are going to
 3
              THE COURT:
    be talking about, things from those spreadsheets?
 4
 5
                                  That's correct.
              MR. TYLER:
                          Yeah.
              THE COURT:
                          How many different images does that come
 6
 7
    in to play on?
 8
              MR. TYLER:
                          So it comes in for figure one and then
 9
    there is an additional spreadsheet which I think is in our
10
    exhibit list as 22, and that one has an additional four figures
11
    in it as well, so spreadsheets associated with four figures.
12
              THE COURT:
                          So 5 out of 13?
              MR. TYLER:
13
                          Correct.
              THE COURT:
                          You are looking at the data?
14
15
              MR. TYLER:
                          Correct.
16
              THE COURT:
                          Again, the two categories I'm familiar
17
    with is this terminal digit issue and the pH scale, but I think
    he talked about some other problems he had with data, but they
18
19
    seemed very sort of targeted.
20
              MR. TYLER:
                          Yeah. The other -- so the other data
21
    thing I think is actually where he referred to the fact that
22
    the 12 and the 13 didn't match the data, that has actually --
23
    comes back to that spreadsheet in Exhibit-22 and that
24
    particular figure is -- has been extracted into this
25
    spreadsheet here in Exhibit-22.
```

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THE COURT:
                          Okay.
                                 I think I understand.
 1
         And then is his proposed opinion on these things that
 2
 3
    these figures are fabricated or that they're different from the
    raw data that they can't be traced back? What do you want him
 4
 5
    to be able to say?
              MR. TYLER: I mean, I think our preference would be
 6
 7
    like he says all of those things, like they are sort of
 8
    building blocks. First he says they match and then, you know,
 9
    they can't be traced and then he says that the -- that it
10
    doesn't -- the research record doesn't match the underlying
11
    data, and then he says, it's fabricated. Our preference would
12
    be to do all of that.
13
              THE COURT: Okay. So I'll come back to you now that
14
    we know what we are dealing with. Let me go to Ms. Beidel
15
    since it's her motion as to what issues she wants to discuss,
    how and if we should be excluding anything here.
16
                          Your Honor, I've been informed that it's
17
              MR. TYLER:
    33 reports tracked back to the 13 -- back to the 13 images.
18
19
                          When you say "33 reports," are they --
              THE COURT:
20
    those cover the opinions, but do those include sort of drafts
21
    of the same report?
22
              MR. TYLER:
                          No.
                               Those are --
23
              THE COURT:
                          Thirty-three final reports?
24
              MR. TYLER:
                          Yes. And that is what was introduced
25
    today was all the final reports. Obviously, we produced the
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Page 159 of 206
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other reports. 1 2 THE COURT: Okay. Your Honor, first of all, with respect 3 MS. BEIDEL: to the number of images and the number of reports, this is 4 5 something we flagged with the motion to dismiss stage that I'll raise again. 6 7 The indictment discusses proposal one and proposal two 8 which deal with, as far as I understand it, only figure one 9 from grant one and figure one from grant five. There have not 10 been any 404(b) notices regarding these other figures. 11 understand the counts, they are about discrete conduct, so in 12 the defense's view, one narrowing that can and should happen is 13 that the government should be presenting evidence about the 14 items that were contained in the indictment as the allegations 15 and not all of these other images, that as far as I know, have only been listed on that preliminary list, but never noticed 16 17 for any purpose through a 404(b) notice or otherwise. So I'm looking at the indictment. 18 THE COURT: 19 references five grants, doesn't it, or not? 20 MS. BEIDEL: It does, but if you look at the language 21 of the counts, in each of the counts, there is a discrete 22 reference to one and only one proposal and between the four 23 counts, there is proposal one and proposal two which gets them

> THE COURT: Are you saying that the -- I don't know

only two figures out of this set of images.

24

25

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how many we are talking about -- the 13 images go beyond one
 1
    and two. Do they also go beyond the five grants that are
 2
 3
    referenced?
              MS. BEIDEL:
                           They do not go beyond the five grants.
 4
 5
              THE COURT:
                          They are all within the five grants.
              MS. BEIDEL:
                           Proposal one and proposal two are
 6
 7
    subsets of grant one and grant five. And then the remainder
 8
    are the rest of grant one and grant five.
 9
              THE COURT:
                          Okav.
10
              MS. BEIDEL:
                           Moving on to the Daubert argument, Your
11
    Honor, the standard for expert testimony, of course,
12
    scientific, technical or specialized knowledge that will help
13
    the trier of fact. The big indicia of that is it based on
14
    sufficient facts and data, are they applying reliable methods
15
    and have they between applied appropriately in this case.
         First on the sufficient facts and data here. Dr. Brookes
16
17
    just isn't in a position to know one way or another whether
    there are other images that Dr. Wang preserved or not.
18
19
    CUNY destroy them or move them? Are there other intervening
20
    factors? How thorough was the government's collection?
21
         A lot of his testimony turns on really the absence of
22
    these images, so without acknowledge as to why that absence
23
    exists, he's drawing conclusions based on an incomplete data
24
    stat, and he does not know the reason for that incompleteness,
25
    and I think that's a fundamental flaw in the data and facts of
```

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all of his opinions are based on.
 1
 2
         As to reliability --
              THE COURT: Before you get to that, let me ask:
 3
    ones he's calling the raw images --
 4
 5
              MS. BEIDEL:
                          Yes.
              THE COURT: -- do you acknowledge that they were, at
 6
 7
    least, things that Dr. Wang generated? Maybe they weren't the
 8
    real raw image, but they are connected to this research in a
 9
    way that it's a comparison is kind of -- not a crazy thing to
10
    do?
11
              MS. BEIDEL:
                           Yes.
12
              THE COURT: You can compare this. You might arque,
13
    well, that's the wrong image, but it's part of a sequence of
14
    images that at least it's a legitimate thing to do.
15
              MS. BEIDEL: We are not contesting the visual
    similarities between the two sets of images that Dr. Brookes is
16
17
    flagging in terms of the background around the white boxes.
                          When you say "similarities," are you
18
              THE COURT:
19
    saying you are not contesting they are similar, or that you are
20
    not contesting that they are identical or clearly not the same,
21
    which seems to be what he's saying?
22
              MS. BEIDEL:
                           We are not contesting that they are
23
    clearly the same in some ways. There is handwriting that is
24
    the same as between some of them, for example. Our position, I
25
    think, is what came out on cross, that there are other images
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25

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that are different exposures or that have been stripped and
 1
    reprobed or some other analysis has been done to them that were
 2
 3
    the digital images that form the basis for the white boxes and
    then that those white boxes were cut and pasted by Dr. Wang on
 4
 5
    top of what Dr. Brookes calls the raw image as a sort of
    Ladenol look.
                   That's how Dr. Wang maintained a record of which
 6
 7
    lanes in particular the white boxes derive from.
 8
         It's important to keep in context that this is all
 9
    clinical research with lots of different participants and
10
    different treatment values, so there is a lot of data to keep
11
    track of. It's not just we are looking at 33 images or 33
12
    reports and 13 images, but he was doing thousands and thousands
13
    of Western blots a year. And so as that image is sort of
14
    developed into the final, it was important to keep it somewhere
15
    that he knew where it derived from. So he put it over that,
    otherwise, useless raw data.
16
                         Otherwise, useless raw data from some
17
              THE COURT:
    other test?
18
19
                           From the same test, but a different
              MS. BEIDEL:
    exposure or a different -- it wasn't stripped and reprobed.
20
    Some kind of different treatment that made that test not the
21
22
    ideal visual representation of data.
23
         So what the exposures do is essentially made the
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representations of the data more palatable for publication.

You might have a super dark one that you can't use. You might

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have a super light one that you can't use. The scientists sort of nuance that exposure time to get to the ideal, and that's what we are talking about here. Yes. It derived from the same data, but it's not the actual original that should be used in Dr. Brookes' analysis and, frankly, I don't think he can say -he testified that he can't rule out that possibility. case where the burden of proof is beyond a reasonable doubt, that kind of opinion just is not helpful to the trier of fact. Moving on to the Daubert factors, there is no indication that this image analysis methodology that he's developed and, frankly, that he says is a young developing field with undefined standards, there is no indication that it's been tested in any kind of thorough way. There is no rate of error we can assign to this kind of methodology as applied to a particular Western blot images. There is no type of peer review process that the government or Dr. Brookes has pointed to in support of his method. What there is is a handful of publications that Dr. Brookes himself authored with no coauthors talking about methods of image analysis, but not exactly the same methods as the ones that are applied here. Just because image analysis exists as a concept somewhere in the scientific community does not mean that this three-stage pipeline with eight other assorted scientific testing methods is the correct way to do image analysis certainly in a criminal

case.

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The government hasn't pointed to a single case where this method has been analyzed or applied. There are some ORI decisions affirming administrative decisions from ORI, but they do not assess at all the substance of the methodology.

THE COURT: So the things I have from the ORI, I think in the record, are just sort of what they did -- what the result of the analysis was. They don't really speak to what the analysis that they went through was, do they?

MS. BEIDEL: Not that I'm aware of, Your Honor. as far as I know, there is not a court anywhere in the country, civil or criminal, that has ever been asked this question before, that has been put to the test of whether this is reliable evidence.

So, for it to be for the first time applied in a criminal proceeding such as this where Dr. Brookes, frankly, gave a lot of opinions about what was the scientific standard that were not based on the records, that were just based on his This is the standard for this, for that or the experience. other thing. That's dangerous to the trier of fact.

So let me ask: He lists about eight THE COURT: different forensic methods. Assuming for the sake of argument I agree with you. Maybe this full methodology has never been tested, never been reviewed. No one has any idea how often errors are made with it. Some of these individual techniques

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whether it's using this Photoshop curves thing or the
    brightness in contrast on -- I don't know if that was on
 2
    PowerPoint or something, Adobe, or using this ORI droplet
 3
    product that ORI itself I think puts out there, do you see any
    problem with using those techniques in the sense that that
    technology is inaccurate for some reason, or are you okay with
 6
 7
    the idea that anyone can do that the way someone can perhaps
 8
    show, put something under a black light or something, or do
 9
    something else to sort of enhance how you look at it. And it's
10
    really more about what conclusions you draw from it than it is
11
    from the use of the technology.
12
              MS. BEIDEL:
                           I don't think we're in a position to
13
    contest the individual techniques. We were looking at this as
    a method. You know, what Dr. Brookes' testimony is, as I
15
    understand it, is that to conduct this analysis as he feels
    comfortable with it it requires all of the steps and all of the
16
    methods, so I'm not sure whether he would say, for example,
    that one, if you took out one method, the histogram analysis,
18
19
    for example, that that would be sufficient or helpful to the
           I don't know the answer to that because that wasn't the
20
    particular way that he framed his opinions here.
22
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What I do know is that there are some AI tools being developed that can be used to either corroborate or not the conclusions here. Some of the relevant --

> I think we are probably a really long way THE COURT:

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    from taking an AI tool and saying that's reliable scientific
 1
    evidence, at least that's how I look at it.
 2
               MS. BEIDEL: Understood, Your Honor.
 3
               THE COURT: But you are saying we don't even have
 4
 5
    that here?
               MS. BEIDEL: Some of the journals in play here, if
 6
 7
    you analyze Dr. Wang's work, use those tools and cleared his
 8
    work.
 9
         So, at a minimum, I think a reliable method of analysis
10
    would take into account that work that was done, consider
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whether, you know, as a factor it should be replicated and then factored into the decision making.

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There is also no forensic analysis conducted here, as Dr. Brookes suggested. There are a lot of cases in the criminal realm, child pornography, for example, that talk about the value of doing MD5 hash analysis to make sure that the images that you are talking about trace through forensically.

As far as I know, the government doesn't have some other witness to do that to. Their case rises and falls on Dr. Brookes, and it's all about Dr. Brookes saying -- I looked at these images. I don't really know how they were derived. don't know if they are a complete set of data or not, but I am reaching the conclusion that they either match or don't match using these tools.

That's all that there is. And I think in other types of

case law like child pornography cases, for example, that would not be enough for a Court to find that there is a forensic match between the images.

on that. So the -- again, these tools, the various tools, some are probably more accepted than others, whether it's the ORI droplets or change the contrast on something. The histogram thing, I'm not sure I've seen that discussed in the literature, but the individual tools, it sounds like your view as well. If someone wants to apply those tools and some of them are commercially available, they can do that. It probably doesn't require -- at least in my view, it may not require sort of peer review of, you know, changing the contrast or something to get something to look right. Maybe not.

But you are concerned that if you look at those things either together or separately and reach the conclusion that it's fabricated or something like that, that's where you say it hasn't been tested to that level of analysis?

MS. BEIDEL: Yes. And adding an expert opinion on top of that. So, for example, if we talk about brightness and contrast I think is a good example.

If the government wanted to go into Photoshop and use brightness and contrast to adjust one of the images to 80 percent, and then put up a slide in closing arguments that compares those two images, and make an argument to the jury

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about whether they match or not and whatever that means, we don't have a problem with that. We could make the counter-argument to that.

My problem is where Dr. Brookes comes in with the imprimatur and the PhD from Cambridge and espouses this method as if it's, you know, you went in to any lab in the country, and you would find that this was a generally accepted method. And he says these images are fabricated. We're replacing a jury's role with his role, frankly. He presents well.

You know, I struggle to see what jury would listen to the way that he presents that with the level of conclusion he's willing to reach about what's accepted and what is not in the scientific community and sort of dare to differ with it, frankly.

THE COURT: So, what if he got up there and said, look, I have looked at these side-by-side documents, and here is some side-by-side images I put together using some of these tools, whether the brightness and contrast, or the color gradient, and so now you can kind of see with this color how similar they look, how different they look. And so here is what I generated through these things and also to some degree I'm going to point out these two things kind of match up on both screens, these things look different, but without reaching the conclusion there is a fabrication. What would be wrong with that?

25

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MS. BEIDEL:
                           I think that's the jury's role, Your
 1
 2
            I'm not sure what is necessary --
                          No. Not saying the fabrication, but just
 3
              THE COURT:
    putting up the side-by-side comparisons.
 4
 5
              MS. BEIDEL:
                           So there is a case Dorsey in the Fourth
    Circuit that talks about comparison of surveillance videos, for
 6
 7
    example. The Court found that the jury's rule was supplanted
 8
    when the expert was essentially just comparing two things with
 9
    each other.
10
              THE COURT:
                          But these are not raw images.
11
    things that had to be enhanced somehow through these tools, so
12
    you would probably need somebody to explain why they look
13
    different, these are different colors. What red means is this
14
    and what orange means is that.
15
              MS. BEIDEL:
                           I think that's a little bit closer to,
    you know, a custodian of a tool sort of evidence.
16
                                                        This is the
17
    kind of -- you know, a police officer talking about how they do
    radar detection or a little closer to that if we are talking
18
19
    about a tool as opposed to this overarching methodology.
                         And then what about his opinion in
20
              THE COURT:
21
    looking at the two side-by-side images which maybe we can have
22
    him put up and say, well, these are the same or different
23
    because I look at the background noise and to me they look the
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same or they look different or they are exactly the same?

I think there is a fundamental flaw in

MS. BEIDEL:

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that opinion where he's not certain that he's comparing the 1 right things. If it's not an apples-to-apples comparison, then 2 that opinion doesn't have that, so if he does not have the 3 correct original, which we -- it's our position that he does 4 5 not -- and it does not matter whether or not the raw image matches the white box image. It does not foreclose the 6 7 argument that we're making which is that the white box was 8 pasted on top. 9 THE COURT: Well, sure. But, again, that goes to --10 you're almost saying he should be able to say that, and we are 11 going to have our other argument. 12 MS. BEIDEL: I understand that goes to the weight in 13 a sense, but I think it also goes to the reliability of the 14 opinion. Because this is such a self-created standard for him, 15 if he cannot foreclose the possibility that he had the wrong original, then it's not a reliable opinion. 16 17 It's very similar to his colleague that he made an allegation against. If that person didn't have that original 18 19 film and come in and tell him he was wrong, he would have 20 persisted in the opinion. That's the case here, too. Dr. Wang 21 should not be made to prove a negative. If these films existed 22 and were destroyed for whatever reason by CUNY or not collected 23 by the government, that's not on him, and I don't think that that makes the method reliable. 24 25 THE COURT: What about the other parts of his

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    opinions or expertise? It seems that there is no dispute that
 1
    he has some expert on using Western blots. He tried to show
 2
    how they work. Any problem with the government having somebody
 3
    do that just so the jury understands what a Western blot is?
 4
 5
              MS. BEIDEL:
                           No, Your Honor. No problem with that.
              THE COURT:
                          And what about his data analysis?
 6
 7
    have this, in some cases it was less about the image and more
 8
    about the data. For example, the pH scale. At one point I
 9
    think he said, well, this doesn't match with the way things
10
    work.
11
              MS. BEIDEL:
                           Our biggest problem with things like the
12
    data analysis, Your Honor, is it seems like a choose your own
13
    adventure method. Whatever image he gets to, he uses whatever
14
    tools he thinks are appropriate in that circumstance.
15
    not some application of that tool in other images to make the
    counter-veiling point that it does not prove that point, that
16
17
    the data is fine, and all the other circumstance, for example,
    or that the histogram analysis only shows that one out of the
18
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So a method generally is a step-by-step analysis requiring conclusions to be reached or not. And this feels like he's just, you know, applying his experience to choose whatever method he thinks is appropriate under the circumstances.

13 images appears to have been manipulated.

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THE COURT: But in terms of the method of saying -and, again, I have to go back and look at the transcript to get

```
the exact opinion, but my sense of the flavor of it was that
 1
    this pH scale is something that comes into play and the data
 2
    does not match with the way it would actually turn out with a
 3
    legitimate scientific experience. And then the same thing with
 4
 5
    this terminal digit analysis he's got. I mean --
              MS. BEIDEL:
                           Yeah. I suppose if those were part of
 6
 7
    generally accepted methods of image analysis, then we wouldn't
 8
    have a problem with it. I'm not so much fighting the substance
 9
    of those things. I'm fighting the conclusion --
10
              THE COURT:
                          Not part of the analysis.
11
              MS. BEIDEL: -- that there is no generally accepted
12
    method for this kind of analysis in the community. He calls
13
    himself -- he's out -- he's operating outside the scope of his
14
    university. There is no peer review process at the university
15
    that looks into that. They make sure he's separate.
    doesn't have any coauthors on any of these papers on this
16
17
    topic, which is pretty unheard of.
         There is people on blogs all over the place either
18
19
    espousing his view or the opposite. And if we bring that level
20
    of discourse in to a criminal courtroom, it really sets a
21
    dangerous precedent.
22
              THE COURT:
                          Okay. Anything else you want to offer at
    this point?
23
24
              MS. BEIDEL: That's it, Your Honor.
                                                   Thank you.
25
              THE COURT:
                          Mr. Tyler?
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A few points I would like to emphasize.
 1
              MR. TYLER:
         First of all, just to -- I'm not sure Your Honor is aware,
 2
 3
    but we engaged with the Daubert factor. If you read Daubert
    itself, it says, many factors will bear on the inquiry as we do
 4
 5
    not presume to set out a definitive checklist or task, but some
    general obligations are appropriate and that --
 6
 7
              THE COURT:
                         Can you please just speak a little slower
 8
    for the court reporter.
 9
              MR. TYLER:
                         Yes.
                                I apologize.
10
         And the Fourth Circuit in the United States versus Crisp,
11
    at 324 F.3d 261 says, as the Court explains -- the Supreme
12
    Court -- the addition of the new factors will put an end to
13
    wholesale exclusion of expert testimony based on scientific
14
    innovations under an uncompromising and general acceptance
15
    test.
         Now, I think what we hear from the defense here is like a
16
17
    broad-based attack, not achieving have --
                          Let me start with some background.
18
              THE COURT:
    mentioned Daubert. Just make sure I'm not missing something.
19
20
         Dr. Brookes has never testified as an expert; is that
21
    correct?
22
              MR. TYLER:
                          That's correct.
23
                          Are you aware of any other expert who's
              THE COURT:
24
    testified in this field in a criminal case or a civil case in
25
    federal court or state court?
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MR. TYLER: No.

THE COURT: So you can't find anybody else, so it's not as if there are other experts who know more. good as it's going to get, it seems?

MR. TYLER: Yes. I mean, that is correct. there is ORI which I was going to say about something about in a second.

THE COURT: On ORI, I wanted to understand what is the state of the record on that because as I looked at it, it seems as if ORI is concerned about this issue. I think they probably don't do a lot of their own investigations. like they set some standards or quidelines because I quess a university is probably supposed to self-police maybe, but then they put out things liking the droplet and say, you can use this to try to figure out what is going on, but they don't really explain how to do it and they don't set any guidelines on sort of how sure you have to be or whether certain tools will show something that clearly shows a misconduct or not. But they give, at least, some tools, at least one of which, maybe more, Dr. Brookes uses, but they don't kind of have a standard methodology that they can say -- that Dr. Brookes He's not following anything that ORI set up, other than perhaps using some of these tools that we have talked about.

> MR. TYLER: Yes. A couple things on that, Your

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also does.

Honor. 1 First of all, just broadly, based on our interview of a 2 3 couple of our people, one in particular, our understanding is you are right. In a lot of cases, ORI directs the university 4 5 to go out and conduct an investigation and they would give a lot of input and advice about ways they may be. 6 7 In some instances, ORI does the investigation themselves 8 if they are not satisfied with what the university or other 9 institution did, and we --So they have to have staff for that? 10 THE COURT: 11 MR. TYLER: Yes. They have ORI investigators. 12 Scientific investigators. 13 One of them, who we interviewed, talked about how they use 14 magnification and comparison, contrast and brightness 15 adjustment, curves functions, gradient maps and densitometry analysis in the work that they do to reach their --16 17 THE COURT: Okay. Hold on a second. I think I jotted those down. I am familiar with those. 18 19 Do they use the other techniques? So do they use the 20 histograms? Do they use terminal digit analysis? Do they use 21 pH scale analysis or things like that? Have they ever done 22 that. 23 I don't know if they have ever done that. MR. TYLER:

The things I gave you are the common things that Dr. Brookes

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THE COURT:
                                 And the droplets, I assume,
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                          Sure.
  because that is kind of their thing.
2
                                And that actually leads to the
3
             MR. TYLER:
                         Yes.
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other point which is we submitted a little over a dozen of Dr. Brookes' reports to ORI during our investigation, frankly, because none of us are scientists, and we wanted to make sure we were understanding things correctly. And they basically This looks good to us, and they gave some feedback about, frankly, doing some more ORI droplets which is what resulted in the final reports.

> THE COURT: Why is nobody from ORI testifying?

MR. TYLER: The short answer on that is that the -the short answer on that is that because that is not who the expert is who we employed. Frankly, at the time we did it, I don't think we had a full understanding of what ORI did.

So the aspect of ORI I had some THE COURT: Okay. uncertainty about was to what extent Dr. Brookes has done work or submitted reports in other matters to ORI, and they have said -- they have sort of validated his results and to what extent any record of that one way or the other. Some of it is have they actually said yes. Your analysis was correct. you did it the right way. And to what extent is it sort of more circumstantial like, well, he submitted something to somebody and it ended up at ORI and then somebody else got suspended, but we don't know exactly how the line runs through

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those things.
 1
                          I think it's somewhere between those
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              MR. TYLER:
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            Dr. Brookes, as I understand it, submitted his
    referral directly to ORI, but I don't know that we have insight
 4
 5
    in to whether they accepted that in total or they did
    additional work themselves, but like what we do know is --
 6
 7
              THE COURT:
                         Submitted something you mean about this
 8
    case or about other cases?
 9
              MR. TYLER:
                         About other cases. And some of that
10
    would be found in the scientific integrity CV if you look at
11
    that. So he submitted like referral information he did to ORI
12
    and ORI ultimately made a finding. I don't know that --
13
              THE COURT: Is there a list there? I think there is
14
    something where -- whether he's provided or someone else that
15
    tells us how often that happened.
                          I think it's just two or three times
16
              MR. TYLER:
    which I believe there is actually a footnote to our response,
17
    if I remember correctly.
18
19
              THE COURT: Are there times where he submitted
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    something and nothing came of it? Either they said, sorry, you
21
    are wrong, or they just didn't react to it in any way that
22
    showed validation of any kind?
23
              MR. TYLER: I don't know the answer to that question.
24
              THE COURT: So, if there is two or three that worked,
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    it may arguably in a circumstantial way showed that they agreed
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with him, we don't know what the total denominator is, how many
 1
    times he did that?
 2
              MR. TYLER: I can't tell you. I haven't asked that
 3
    question. I can't tell you that.
 4
 5
                          I take it ORI probably doesn't have that
              THE COURT:
    kind of data, like they are not tracking that kind of thing?
 6
 7
              MR. TYLER:
                          I don't know that for a fact, but my
 8
    impression is no.
 9
                          And then in terms of the literature we
              THE COURT:
10
    have seen, it seems to me that there are articles that cover
11
    this area or at least talk about it, less about doing an
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    experiment, and I think, as Dr. Brookes said, a few of them are
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    just, well, this is a problem, and we tried to figure out how
14
    bad it is, and they use their own methodology to figure out how
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    often people engage in misconduct and came up with some numbers
    sometimes using either an AI tool or this I think he said he
16
17
    named something that came out of one of those like a
    proprietary software that he said he can't get access to now.
18
19
              MR. TYLER:
                          A C lot.
                          C lot. Or one said he had like three
20
              THE COURT:
21
    scientists looking at it together and deciding if they all
22
    agreed. So they all had different ways of doing it to identify
23
    the problem.
24
         Is it fair to say that nobody has sort of used this
25
    three-step process that's written about it that Dr. Brookes is
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    using here?
 1
                          Not -- I don't think not specifically.
 2
              MR. TYLER:
 3
    think his testimony actually is the best which is like this
    three-step thing is really just about figuring out if A goes to
 4
 5
       Can the published image be tracked back to a raw image?
    And like he laid out this way of doing it, but really that is
 6
 7
    what really all of these articles are asking the question.
 8
    you using similar tools and different -- like different ways of
 9
    doing it, but that's the question that they're all talking or
10
    using or all asking and they are all using at least some of the
11
    same or similar tools.
12
              THE COURT: So, other than using the same tools,
13
    maybe some of which might be generally accepted, there is no
14
    real general acceptance of his approach or methodology;
15
    correct?
16
              MR. TYLER:
                         Not in like -- not framed in like
17
    three-stage standard.
                          In terms of whether it's been published
18
              THE COURT:
19
    in any way, I think the article I think that's closest is this
20
    detection article, that he -- misconduct detection article.
21
    don't think it goes through the three steps. I think it lists
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many of the technique, if that's right.

MR. TYLER: Yeah. That's my understanding, Your

Honor.

25

THE COURT: And as he described that he spoke and

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they said everybody can submit an article. So his list of 1 techniques is published. Was that -- do you know if that's a 2 peer review process to be published for that or given the 3 symposium nature of it, it wasn't? 4 5 MR. TYLER: That is a fair question. I don't want to answer without -- I don't know for sure the answer to that. 6 So 7 I don't want to tell you the wrong answer. 8 THE COURT: Sure. I mean, I was inferring from the 9 way that you described it there was not necessarily a peer 10 review. But do you have any reason to or any information that 11 contradicts that in some way. 12 MR. TYLER: No. I don't know that. 13 THE COURT: So I think where you started on Daubert 14 you were saying -- and I think I agree with this -- and Kumho 15 Tire, too. There are these five factors. None of them are dispositive. We are probably supposed to look at all of them 16 17 when there is any kind of technical stuff going on which is the case here, so we will look at all of them. 18 19 We are not limited to those, but the five of them don't --20 I mean, given that he's acknowledged it's a new field, they 21 don't really map on very easily to say, well, these five 22 factors all work so his methodology is totally set on. 23 For example, error rates and testing. This seems like the

kind of thing someone could test. You could get some images,

either fabricate them or just take existing images and run them

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through somebody and say, hey, can you tell me if these are the
 1
    same or different, and you know the answer.
 2
 3
         That is kind of happens with handwriting in the Crisp
           They do testing, both broadly to see if their
 4
 5
    methodology works, and also the individual examiner.
         None of that has ever happened here; correct?
 6
 7
              MR. TYLER:
                         I mean, you -- I think you cited a Bik
 8
    case where she went through with some other co-authors and they
 9
    identified as to the --
10
              THE COURT:
                         I'm sorry. Say that again.
11
              MR. TYLER:
                         The Bik article which is one of the ones
12
    where she goes through and like identifies, but it's not --
13
    it's really the flip of that which is say we have identified
14
    this number by this process, but not the error rate of the
15
    actual process.
16
              THE COURT:
                          Is this the one where there are three
17
    examiners looking at it together?
18
              MR. TYLER:
                          Yes.
19
              THE COURT:
                          But that's a different methodology?
              MR. TYLER:
20
                          Yes.
              THE COURT: So he has not even done it himself.
21
22
    hasn't like tested it himself and said, hey, someone give me
23
    all these images, and I'll tell you if they're fabricated or
24
    not and see how I do.
25
              MR. TYLER:
                          I think that's right.
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THE COURT:
                          So, if the traditional factors, at least
 1
    don't obviously map on to the idea that this should be
 2
    considered reliable, what would you point to, again, since we
 3
    are not limited to those five factors, but given that it has
 4
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    not been tested, and there is no error rate for his work, there
    is no -- or the methodology is not generally accepted.
 6
 7
    would you point to to say, nevertheless it's reliable?
 8
              MR. TYLER:
                         A couple things. One, I think you could
 9
    argue that the error rate is one instance. And I think he may
10
    have said it in other sentences that we disclosed that he
11
    estimates that would have been a percent or something of the
12
    overall number of situations where he's alleged misconduct
13
    based on the process that involved.
14
                          I'm sorry. What did you say about
              THE COURT:
15
                  How did he get there?
    one percent?
                          So he talked about the one instance where
16
              MR. TYLER:
    he was confronted so, obviously, he admitted that was an error
17
    there and like none of the other instances where he's
18
19
    confronted has that happened to --
              THE COURT: I'm not sure he said that. I think he
20
21
    just --
22
              MR. TYLER:
                          Yes. But the point is like he's got all
23
    these other papers, like hundreds of papers retracted, and this
24
    is one instance.
25
              THE COURT:
                          Where does it say hundreds of papers?
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didn't hear him say that today.
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That's in his -- that's actually in his MR. TYLER: He lists some of them. CV.

Again, maybe I missed it, but the same THE COURT: question as I had with ORI. Do we know what the denominator If he's gotten a hundred retracted -- just as an example. Let's pick a number of a hundred. How many has he submitted something where there was no retraction? Do we know?

> MR. TYLER: I don't know that, Your Honor.

If I may, though, to answer your original question is like -- is, frankly, part of what Ms. Beidel said which is like we are not even contesting that these things match. lot of what the point of today was from our perspective was to actually like show the work, demonstrate the reliability of this using multiple tests with tools that Dr. Brookes did not invent and that are generally accepted and available. using those tools like walk you through why this is reliable.

The defense's argument is all like a broadside against this idea of like what are the standards without actually engaging with like, there is something unreliable about this conclusion he reached or about this analysis he did. And the reason for that is because, notwithstanding the fact that like it would be hard for a juror to understand what was going on in looking at this cold, but like you can actually see it and it is apparent.

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like --

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          THE COURT:
                      Let me ask this:
                                         A couple times he said
there is a digital fingerprint, and that caught my attention
because, as you know, fingerprint analysis is under fire now.
It used to be everybody thought these were exactly the same.
And they have a lot of standards. They have five points of
comparison that need to be matched between two different
fingerprints and even then, if we don't have absolute
certainty. He said just by looking at the background noise, as
he called it on these two, he could tell they are exactly the
same or different. There is a digital fingerprint there and so
he is certain that this slide came from that one.
     I understand the notion of saying, well, there is a common
sense -- it makes sense to me if he says as someone who uses
Western blots that if you go back and try to use the same thing
twice, it is not going to have the same background noise.
even that, when we have years and years of fingerprint
analysis, it is not certain that two people don't have the same
fingerprint.
     How can you be so certain about that? What is he basing
that on?
                      I think in that instance when he's
          MR. TYLER:
talking about the matching background, the point is instead of
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THE COURT: And we don't have five points of

comparison between the two backgrounds. I just have a fuzzy

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image on a screen, and I'm taking his word for it that they're 1 exactly the same even more so than two fingerprints are exactly 2 3 the same. I think what he would say is he would MR. TYLER: 4 5 point you to there is way more than five commonalities here 6 that I can point you to, and these are the ones that I looked 7 at when I made that comparison. 8 THE COURT: But what's the standard for how many you 9 have to match to say these are exactly the same? 10 MR. TYLER: I mean, I think his testimony governs it 11 right there. I don't think we can point to --12 THE COURT: So how is this more reliable than 13 fingerprint evidence, which is in question, or handwriting 14 analysis where you compare different parts of handwriting which 15 has been vastly tested? This has never been tested. 16 just what he's saying. 17 MR. TYLER: I think the answer to that is the difference between fingerprint analysis is really the 18 19 fingerprint analysis is you are comparing people's -- everybody 20 has like a bunch of fingerprints. And you are just really 21 trying to like figure out like this line versus that line. 22 Right. 23 Here you have like the things -- whatever are the 24 fingerprints here are not necessarily -- especially if you are 25 talking about the background artifacts, like they like don't

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all have to be here. The fact that they are emerging in the --
 1
    in the way that they are with the -- you know, with the
 2
 3
    orientation, with spacing, with the size and shape, like that
    like just --
 4
 5
              THE COURT:
                         How do we know they don't just have a
    sort of a particular thing that's stuck on the glass somewhere?
 6
 7
    How do we know these things? How is he so sure?
 8
              MR. TYLER:
                          I mean, I think that the answer to that
 9
    is based on the experience that he's been doing this for 20
10
    years, that he knows that this is how this works.
11
              THE COURT:
                         Does anybody else do that analysis?
12
    didn't see any other articles where there was a discussion not
13
    about how you look at the bars or the bands, but the background
    noise piece. I didn't see that as a discussion point of here
14
15
    is how you tell the background noise tells you these are the
16
    exact same things.
17
         Does anybody else do that, other than Dr. Brookes?
                          I am not aware of -- I'm not aware.
18
              MR. TYLER:
              THE COURT:
19
                          Does ORI use that type of analysis?
20
    Does ORI have any standards for that?
                          Not that I know of.
21
              MR. TYLER:
22
         Your Honor, just to also close a loop on that. Obviously,
23
    it's not just the -- you've got the handwriting. You've got
24
    the file name. There is like more than just one data point
25
    here, so we think that all of the -- like, obviously, that is
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part of this as well.

THE COURT: Okay. So --

3 MR. TYLER: Your Honor, if I could point out one

4 other thing, which I --

THE COURT: Sure.

MR. TYLER: -- thought would posit just on sort of looking at this through a different rubric. I mean, 702 and Kumho Tire both explicitly talk about experience being a basis like on its own.

In the notes on 702 it says, nothing in this amendment is intended to suggest that experience alone, or experience in conjunction with other knowledge, skill, training or education may not provide a sufficient foundation for expert testimony.

To the contrary, the text of Rule 702 expressly contemplates that an expert may be qualified on the basis of experience. In certain fields, experience is the predominant, if not sole basis, for a great deal of reliable expert testimony.

And then *Kumho Tire* it says, no one denies that an expert might draw a conclusion from a set of observations based on extensive and specialized experience.

THE COURT: So a lot of times that's like our drug agents who know the lingo. This does seem like we are not far off from the *Kumho Tire* situation where there is an element of science involved. Clearly, I don't think anybody off the

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street could do what Dr. Brookes is doing. I disagree that
this is like the photograph case because you have to know
something about these Western blots to do some of this.
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But I think they had an engineer in Kumho Tire and it's unclear to me whether, you know, just saying I've looked at these things a lot is enough in that context.

So is there a case that you think is a good factual comparator for our situation where it's perhaps not entirely pure science, but is also not just someone saying, look, I've heard a lot of drug dealers talk, and this is how they talk.

MR. TYLER: I mean, I'm -- I think that the -frankly, I think that the handwriting cases are probably the closest. I don't -- like -- as I said before, I don't think there is any direct analysis in this case here.

I think the handwriting case probably is the closest because there is a degree of reliability to it, but there is a degree of subjectivity.

In this case, in my view, it is even more objective here because it's not like, are the things like leaned over this much, but it's more based on like this program and this tool is doing this and making this assessment, and I'm just like reading what the results of that are.

There is no tool being involved with this THE COURT: background noise issue, though. Right? It is really just the ones that try to overlay the bands from the tests; correct?

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Yeah. I mean, he said that -- he MR. TYLER: apologized that he did not do the ORI droplet, but like conceivably, you could do an ORI droplet on that background and get to some sort of result. Obviously, that's not part of what we posed here, but I quess that would be one way to do it. THE COURT: What about the data issues that we heard about the pH scale, the terminology digits? I mean, I think because we are not -- I think part of the nature of the hearing didn't lend itself to sort of challenging every single piece of evidence. What's the basis for the reliability of those pieces of information? I mean, I think the pH scale one is just the fact that he's -- this is like why his expertise is particularly important -- because he's able to say the way you run this, you run top to bottom. And if you look at what they have done, if you run it top to bottom, the bottom is now zero, which is an impossible pH -- an impossible pH level. I, frankly, didn't fully understand it THE COURT: because there was a step in the middle called the vertical stretch, and I didn't understand what that was. I also didn't understand whether that made a difference. MR. TYLER: The vertical stretch was just matching what he was doing in the middle with the same scale of the published image and make sure they were oriented in the same

way so you could actually transfer over in the 3.95 in to 9.5.

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That's an example where his point is that like -- like I
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    know this because I'm a scientist and you have to run these
 2
 3
    things top to bottom, but the problem with this is if you
    actually look, the bottom is like not a -- is like a pH scale.
 4
 5
    It is not possible.
              THE COURT: I guess I would have to go back and read
 6
 7
    it again.
             I guess I didn't fully get it the first time through
 8
    how we know that we are below the scale in some form.
 9
              MR. TYLER:
                          Your Honor, also, all the stuff that we
10
    went over is like in -- so if we are talking about Exhibit-10A,
11
    Exhibit-10 is the report where he actually like gives a
12
    narrative explanation for that, so that's also -- that's not
13
    only just the testimony, but that's also a place where he's
14
    explaining it.
15
              THE COURT:
                          Okay. So, other than the handwriting
    comparison that seems as if just in terms of other indicia of
16
17
    reliability beyond just the five factors, it's more just it
    sounded good. Is that basically the analysis, like he got up
18
19
    there and it makes sense? I don't disagree a lot of it makes
20
    sense, but is that really -- is that what I'm supposed to go on
21
    or --
22
              MR. TYLER:
                          I would not characterize it as that.
23
    just like he's showing the actual step and that was the purpose
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of that initial PowerPoint is just like I'm showing you the

steps that I have done. This is like a reliable software

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And this is what we get here and what this is what we
 1
    get here. And all I'm doing is I'm running those --
 2
    objectively running those programs, and then the result of that
 3
    is just going to be what the comparison is, and then that is
 4
 5
    what it is.
         The matching -- I mean, the version forward analysis, like
 6
 7
    when you actually like do the step-by-step process, it's
 8
    reliable, and it's also reinforcing because you have the
 9
    gradient analysis which matches like the magnification and
10
    orientation that matches the droplets.
11
         So in those instances, you have multiple ways that is
12
    actually like corroborating or reinforcing the reliability when
13
    you are looking at the actual analysis being done.
14
                          So, even if we take the view that some of
              THE COURT:
15
    these steps, like I was telling Ms. Beidel, you are just going
    to put up the gradient image, it's easier to tell what is going
16
17
    on from the colors. Someone can accept the notion that there
    is nothing sort of -- the technology is not sort of changing
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19
         It is just making it easier to see, and Dr. Brookes can
20
    say it's easier to see.
21
         How do you get from there to, "in my expert opinion, this
22
    has been fabricated?" Again, to do that kind of opinion I
23
    think usually the standard for, again, how reliable it is,
    probably should be higher. And I'm not sure, frankly, in any
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of these circumstantial instances we have, whether it's ORIs,

does.

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debarring somebody or somebody withdrawing their paper, that we
 1
    really have any examples where there is a through line from
 2
    using these techniques to finding that something has actually
 3
    been fabricated.
 4
 5
                          I mean, it's a different standard, but
              MR. TYLER:
    overall, if you look at their actual finding, it's a number of
 6
    them find that there has been falsification or fabrication.
 7
    You have a different standard here.
 8
 9
              THE COURT: So the background on those is not
10
    available to anybody? Like the details of those -- again, all
11
    I have is the results. I don't know what methodology they
12
    reached to reach that conclusion. I don't have what their
13
    standard of review is, how high the bar is as a matter of
14
    evidence. Do they use the same techniques that -- I mean, Dr.
15
    Brookes is only using not all of his A techniques anyways.
    Which ones did they use? Did they use anything about terminal
16
17
    digit analysis? I mean, I don't really know. Is that written
    down anywhere?
18
19
              MR. TYLER: I mean, I think some of -- the short
20
    answer -- the thing that I think comes to mind right now is we
21
    have -- like during the course of this, we have like a draft
    declaration that was submitted to an ALG and one of the judges.
22
23
    It's not signed. I don't know if it's final or not, but it
24
    does go through a lot of the same analysis that Dr. Brookes
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There is no terminal digit analysis in there, but it

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talks about using the ORI droplets in current files and there 1 is a formal declaration that's done by the actual ORI 2 3 investigator.

THE COURT: Does Dr. Brookes claim any expertise on terminal digit analysis because it is not -- it's not -- it seems like it makes sense. At the same time I don't know if it's appropriate to say you are just going to do a standard sort of -- you know, he's not a statistician. He decided what he thought was the relevant statistical analysis. I don't know if that is legitimate or generally accepted or anything like that.

MR. TYLER: I think that's fair. I mean, obviously, like you can tell, again, just by going through the spreadsheets, yes, he has a lot of statistical knowledge and he did have graduate statistical training in order to just do what he does, but he's not a statistician, which is true.

THE COURT: Let me ask you this: I have to -- I'm going to have to think about this. One scenario, though, might be along the lines of, as I said with Ms. Beidel, maybe he gets to show the work he did in comparing some of these documents using these tools which are somewhat uncontroversial, like the ORI droplets and putting up images about how this is what the side by side looks like. Maybe pointing out things based on his knowledge more as a user of Western blots as to what some of the similarities or differences might mean from a scientific

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standpoint, but maybe perhaps saying, well, there are certain
 1
    opinions you may be able to give and certain maybe you can't
 2
    because they're just not -- methodology doesn't lend itself to
 3
    saying, for example, this is entirely fabricated.
 4
 5
         What I'm concerned about if I were to do something like
    that is, from his testimony, he just seems to be very confident
 6
 7
    in his views. And if I said, look, you can't call something a
 8
    digital fingerprint, he might actually just do that anyways.
 9
    That's the impression I get about him. Or if I say don't say
    it's fabricated, he might just do that anyways.
10
11
         What controls would there be on a situation like that, or
12
    is that just not helpful to you? Would you just rather have
13
    all or nothing with him?
14
                          We would rather have an all or nothing.
              MR. TYLER:
15
    I do think that we can -- and, frankly, obviously, we prepared
    him in advance of today. And I do think that we can absolutely
16
17
    cabin this as necessary consistent with the Court's order.
                          Okay. Any other arguments you want to
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              THE COURT:
19
    offer at this point?
              MR. TYLER: Can I have a moment?
20
21
              THE COURT:
                          Uh-hum.
22
              MR. TYLER:
                          I don't think anything else, Your Honor.
23
    I think I made my points, unless you have any -- we
    obviously -- I'm happy to address like any more specific
24
25
    comments, but I think the most -- from our perspective, the
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The rest

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most important piece here is like the actual analysis itself
 1
    and like walking through that is what shows it's reliable.
 2
 3
              THE COURT:
                          Isn't that a sort of chicken or the egg
    thing?
            I mean --
 4
 5
                          I don't know about that because like if
              MR. TYLER:
    you say like this handwriting matches whatever. And you are
 6
 7
    just like, that's just like an opinion based on like, oh, I
 8
    have done some training and this is the standard or whatever
 9
    versus like I'm going to show you how this stuff actually
10
    works, and I'm going to show you how this stuff matches.
11
    That -- I'm not going to ask you to take my word for it.
12
    going to -- I -- and I did today demonstrate to you like how
13
    this is reliable and how you can actually see that there are
14
    multiple ways to verify that this is reliable.
15
                         So I understand that you can do a side by
              THE COURT:
    side just as you can with handwriting, and you can point out
16
17
    where there is things look the same and things that don't look
    the same, and the handwriting expert can then, at least if
18
19
    they're admitted -- and I don't think it's a universal thing,
    but it's -- they can say, you know, based on the way on these
20
21
    various points of comparison, I think it's the same person.
22
    I'm not totally sure whether, again, on testing on that they
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have done, you know, a lot of different work not just on the

technique, but also on the individual person as opposed to

just -- so there is that part where it makes sense.

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    of it I have to think about.
 1
         The question I had was: I read the defense's
 2
 3
    submission -- and, again, I'm not saying we are resolving this
    all at the same time, but we might. They have their two
 4
 5
    experts. It seemed as if, first of all, they don't need them
    if they don't have Dr. Brookes here. But if they have Dr.
 6
 7
    Brookes here, then they -- it seemed like your arguments were
 8
    more that they were not relevant. And I think, for example,
 9
    you say that there will be no dispute that meta data does not
10
    trace back, although Dr. Brookes does not have anything to say
11
    on that one way or the other. He didn't try to do that.
12
         So, if I agree with you that, even if it's undisputed, I
    mean, if they want to make the point that there is no meta data
13
14
    or forensic way to tie these things together, other than I
15
    think the main connection is Dr. Brookes saying the background
    looks the same, ergo they are the same.
16
17
         How would they get that into evidence or how could they
    argue that if they don't have somebody saying -- because he
18
    won't be able to say that one way or the other?
19
20
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Well, I think he did say that is that I MR. TYLER: didn't do any of this. I think is the point that they are trying to make is that like not -- frankly, it's probably a better point for them which is that like not only did they not do it, but he like didn't even consider or do it all. Right.

So, I mean, I guess I think that that comes in.

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THE COURT:
             Okay.
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MR. TYLER: With also one other thing is like, Your Honor should have probably picked up on this some, but like some of this testimony is going to connect up with what Special Agent Weeks testifies about what he saw in the files and what like, you know, some of the like time stamps and whatever that were associated with the actual underlying files that were provided to Dr. Brookes.

So there will be some connection in terms of that, but, obviously, Dr. Brookes only has what he's given.

THE COURT: Okay. And then the other expert was about a literature search and on the one hand -- I mean, I think it's specialized enough that it's not -- an ordinary juror would not know where to look, I think, or how to read anything that they found, so you are sort of saying maybe it does not really matter or maybe it does because you said, well, there are articles that he didn't find that are relevant, but they are some of the same ones we have talked about.

Dr. Brookes seems to give the view that, well, you know, it's all out there. Everybody is doing it. And what is wrong with somebody saying, well, I don't really see any articles that talk about this methodology in the same way.

MR. TYLER: I mean, I think that that's -- I think that one of the issues with that is that he reached that opinion before he even reviewed Dr. Brookes' methodology.

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So I think like to reach that conclusion without
 1
    describing what you -- providing no basis for what you have
 2
 3
    done, to reach that conclusion, and then haven't even reviewed
    his methodology is like pretty indicative of a deficient
 4
 5
    methodology in the first place.
              THE COURT:
                                 That's fair. Anything else on any
 6
                          Okay.
 7
    of the experts?
 8
              MR. TYLER:
                          Unless you have any questions, like I
 9
    think our papers stand for themselves on both those issues.
10
              THE COURT:
                          Okay. I mean, I do think that they're --
11
    I'm hoping by going back it will be more clear exactly what his
12
    testimony will be because I think that, to the extent that I
13
    allow it, if with certain kinds of restrictions, it's kind of
14
    hard to do that without knowing the full scope, so we may come
15
    back for more clarification if it gets that far.
         Ms. Beidel, anything else you want to offer?
16
17
    are --
                          Your Honor, just to --
18
              MR. TYLER:
19
              THE COURT:
                          I told the other case that we would start
20
    at 3:00, so I don't want to run too much longer. We gave you
21
    an extra half hour, but if there is anything you want to
22
    respond to quickly.
23
         Was there one last point, Mr. Tyler?
24
              MR. TYLER:
                          I just wanted to point out that like when
25
    you actually look at the reports, most of it is -- most of the
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reports is like that forward and reverse analysis, so that's 1 like the core of it. Even though we covered a lot of other 2 3 stuff, like that is the feature that is most repeated across the rules. 4 5 THE COURT: I see. Thank you. MS. BEIDEL: Just on that last point with respect to 6 7 Dr. Cheaito, Your Honor, he did review the substantive reports 8 at the time that he issued the opinion which contains the 9 forward analysis and reverse analysis. And we told him that was the method. 10 11 So, after the government pointed that out, we made sure he 12 got reports one through three for completeness, but he 13 understood the analysis and the methodology at the time that he 14 issued his opinions. 15 THE COURT: Okay. Do you need those two experts -if Dr. Brookes gets to testify, do all of you want them or does 16 17 it depend on what his opinions or what he's allowed to testify to and what he's not allowed to testify to? 18 19 MS. BEIDEL: Our view is if he's allowed to testify, 20 we do think we would need the experts to talk about the 21 methodology. I spes -- I can't envision exactly what the Court 22 would allow. Certainly if there was a different methodology or subset 23 24 of methodology being applied, then we would talk to Dr. Cheaito

about whether he thought that was acceptable and his opinion

would change. 1

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The meta data analysis we do think is critical, so going to this point of the digital footprint, there is usually a digital fingerprint. It's in the meta data, this MD5 hash value and it doesn't exist.

So, especially given the potential possibility that Dr. Brookes would go beyond whatever limitations are imposed, we do think that is necessary for our rebuttal, if nothing else.

One other thing, I did want to flag on the ORI denominator and I guess the other denominator about retracted papers. our view, that's Brady. If that evidence exists within the government about issues with Dr. Brookes' reports, if there is ones that ORI turned down, we think that we are entitled to see that material prior to trial.

THE COURT: Okay. I mean, that's a good point. don't know if I -- it may or may not be Brady, but I certainly think it would be material to the defense to know that. either way, it sounds like you don't have that information; is that correct, Mr. Tyler?

> That's correct. MR. TYLER:

THE COURT: Okay. So, as you can tell, I need a little time to think about this and review what we heard today from Dr. Brookes and what your responses were to the questions. And, you know, it's actually kind of unusual to have a proposed expert who's, A, never been an expert. Usually they come in

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and say, "I've done this 50 times," or someone who is also in a
 1
    relatively new field.
 2
         So, forgive me if I need a little more time on it.
 3
         The only thing I want to try to figure out is what is the
 4
 5
    timeline between now and the trial. I think we have one motion
    in limine which is not unrelated to all this.
 6
 7
         I think the response date for that and for the jury
 8
    instructions I think is tomorrow or something like that.
 9
    Usually I try to sort out motions in limine and jury
10
    instruction disputes, if there are any. We hope it will be
11
    jointly agreed to. And we discuss voir dire questions and the
12
    like at the pretrial conference itself.
13
         I guess one question is: Does it present problems for the
14
    parties if the ruling on the expert motions in limine doesn't
15
    come until the pretrial conference? Is there some compelling
    need to have an answer before that?
16
17
              MR. TYLER:
                          Your Honor, not from the government's
    perspective. I think our intention is to prepare. Obviously,
18
19
    I mean, one benefit of today is we did, obviously, a lot of
    preparation of Dr. Brookes already.
20
                          That's right.
21
              THE COURT:
22
              MR. TYLER:
                          And like, obviously, we didn't go through
    all the reports, but we got like a model of what it would look
23
    like and so we will continue to prepare in that fashion.
24
25
              THE COURT:
                         Ms. Beidel, any issues with that?
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24

25

preference.

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We can make that work, Your Honor.
 1
              MS. BEIDEL:
              THE COURT:
                          I mean, I would like to try to resolve
 2
 3
    this earlier in part because there is these other issues to
    sort out so, you know, if I can do it sooner than that, I'll
 4
 5
    let you know, but I also wanted to see what problems might be
    created by that.
 6
 7
         And then in terms of -- I take it the government did not
 8
    file any motions in limine; correct?
 9
              MR. TYLER:
                          That's correct, Your Honor.
10
              THE COURT:
                          So we will have that one motion.
                                                             We will
11
    have the jury instructions which, again, I hope will be agreed
12
    to, voir dire, verdict forms.
13
         Any other issues that the parties want to discuss at the
    pretrial conference, I do -- while we are here, is there
14
15
    anything else that it would be better for me to know about now
    than at the pretrial conference for playing purposes?
16
            MR. SRIDHARAN: I think one thing, Your Honor, on the
17
    jury instructions, we are working through our various positions
18
19
    on them. A lot of them relate to the same pattern involved
20
    sending the indictment language back. And there is, you
    know -- so a lot of our back and forth are about kind of the
21
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About sending the indictment back? THE COURT:

scope of that and, obviously, every courtroom is different on

that, so I think it would be helpful to know Your Honor's

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MR. SRIDHARAN:
                              Indictment language in the jury
 1
    instructions or whether Your Honor sends the indictment back.
 2
 3
    Generally, you know, so I think that would kind of help
    shortcut some of the --
 4
 5
                         Have you talked to the U.S. Attorney's
              THE COURT:
    Office about this? You have local counsel.
 6
 7
              MR. SRIDHARAN: We will reach out to the U.S.
 8
    Attorney's office and see.
 9
              THE COURT:
                          I think, generally speaking, I don't have
    any hard and fast rules. What I generally understood is that
10
11
    many times one side or the other, oftentimes the defense, would
12
    prefer not to have the full indictment go to the jury.
13
    don't -- not always. Sometimes.
14
         But my general approach is, I don't send the full
15
    indictment back as a matter of course. If one side objects,
    I'm less likely to do so.
16
17
         But two things can come into play. One is that some of
    the language, like the very core language of the indictment,
18
19
    not sort of descriptive language, but just as an example
20
    looking at this, there is paragraph 39 on false statements,
21
    many times that will be -- that language will be in the jury
22
    instructions when we are introducing the false statement
23
    charge, so that the parties -- the jury knows the exact date
24
    we're talking about. It's basically just the statutory
25
    language and I probably would include the statutory language
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1 anyway. So a paragraph like that usually shows up. Not always. 2 In saying that -- you know, and the same would be true of like 3 a wire fraud charge we might have some of the language from 4 5 paragraph 34, and certainly some of the language from Count Two or Three just describing what the wire is. 6 7 But the full indictment as a full document or even the 8 full text, ordinarily I would not include. One exception is if 9 particularly in a conspiracy case, which I don't think this is, 10 but sometimes there is other things where it's material for 11 them to know. For example, what are the overt acts alleged. 12 So, then, that might be reason to send the whole 13 indictment back which we sometimes do, or if one side would 14 prefer doing it a different way, we can sometimes give the jury 15 sort of a list of overt acts which is basically just a cut and paste from the indictment, but it's not called the indictment. 16 17 It's just saying these are the things -- the overt acts have 18 been charged. Sometimes we take subparts and use them because 19 they are material. 20 There are sometimes when both sides agree, we will send 21 the whole indictment back. So there is a range. That is 22 generally how I have seen it. 23 MR. SRIDHARAN: I appreciate that, Your Honor, I 24 think the method you described is what we just did in Judge

Xinis' courtroom for a trial I did last year, so that is what I

```
was planning on proposing, but I --
 1
              THE COURT:
                          Which one?
 2
              MR. SRIDHARAN: About including the charging
 3
    language.
 4
 5
              THE COURT:
                          Excerpts, but not the full indictment.
              MR. SRIDHARAN: Excerpts, but not the full
 6
 7
    indictment.
 8
              And that we also did have conspiracy, so that we did
 9
    copy in the overt acts, but that's not the issue here.
              THE COURT:
10
                          Anything else that would be helpful for
11
    me to know or you to know before the pretrial conference?
12
              MS. BEIDEL: No, Your Honor.
                                            Thank you.
13
                          Nothing from the government.
              MR. TYLER:
14
              THE COURT:
                          Okay. And in the meantime, you know, you
15
    may get questions from Ms. Solomon for our planning purposes.
    Please try to respond as promptly and thoroughly as possible
16
17
    and, conversely, if there is some logistical issue that you
    need guidance on, you can try asking her. I'm not promising it
18
19
    will be an appropriate thing for her to have to respond to, or
    certainly file something in advance along with everything else.
20
21
         So we will get back to you as soon as we can on this
22
            Thank you for the argument and the testimony of the
23
    witness.
              It's been very helpful. Thank you.
              DEPUTY CLERK: All rise. This Honorable Court now
24
25
    stands in recess.
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(Proceedings concluded at 3:10 p.m.) CERTIFICATE I, KIMBERLY A. BURSNER, Federal Official Court Reporter in and for the United States District Court for the District of Maryland, do hereby certify, pursuant to 28 U.S.C. \$753, that the foregoing is a true and correct transcript of the stenographically-reported proceedings held in the above-entitled matter and the transcript page format is in conformance with the regulations of the Judicial Conference of the United States. Kimberly A. Bursner Registered Professional Reporter & Federal Official Court Reporter